=> fil capl; d que 11; d que 15; d que 125
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FILE COVERS 1907 - 3 Feb 2006 VOL 144 ISS 7 FILE LAST UPDATED: 2 Feb 2006 (20060202/ED)

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'OBI' IS DEFAULT SEARCH FIELD FOR 'CAPLUS' FILE

L1 1 SEA FILE=CAPLUS ABB=ON US2003-630143/AP

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L3 110 SEA FILE=CAPLUS ABB=ON GAULT R?/AU
L4 339 SEA FILE=CAPLUS ABB=ON JORDAN F?/AU
L5 3 SEA FILE=CAPLUS ABB=ON L2 AND L3 AND L4
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L4
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L7
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L8
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L18
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L19
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L25 5 SEA FILE=CAPLUS ABB=ON L16 AND ((L17 OR L18 OR L19 OR L20 OR L21 OR L22 OR L23 OR L24) OR (L13 OR L14 OR L15))

=> s l1 or l5 or l25

L177 5 L1 OR L5 OR L25

=> fil medl; d que 146; d que 152

FILE 'MEDLINE' ENTERED AT 13:05:46 ON 03 FEB 2006

FILE LAST UPDATED: 2 FEB 2006 (20060202/UP). FILE COVERS 1950 TO DATE.

On December 11, 2005, the 2006 MeSH terms were loaded.

The MEDLINE reload for 2006 will soon be available. For details on the 2005 reload, enter HELP RLOAD at an arrow promt (=>). See also:

http://www.nlm.nih.gov/mesh/

http://www.nlm.nih.gov/pubs/techbull/nd04/nd04 mesh.html

http://www.nlm.nih.gov/pubs/techbull/nd05/nd05\_med\_data\_changes.html http://www.nlm.nih.gov/pubs/techbull/nd05/nd05\_2006\_MeSH.html

OLDMEDLINE is covered back to 1950.

MEDLINE thesauri in the /CN, /CT, and /MN fields incorporate the MeSH 2006 vocabulary.

This file contains CAS Registry Numbers for easy and accurate

L43	383	SEA	FILE=MEDLINE	ABB=ON	HUNTER K?/AU
L44	19	SEA	FILE=MEDLINE	ABB=ON	GAULT R?/AU
L45	301	SEA	FILE=MEDLINE	ABB=ON	JORDAN F?/AU
L46	2	SEA	FILE=MEDLINE	ABB=ON	L44 AND (L43 OR L45)

L43	383	SEA	FILE=MEDLINE	ABB=ON	HUNTER K?/AU
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L45	301	SEA	FILE=MEDLINE	ABB=ON	JORDAN F?/AU
L51	3885	SEA	FILE=MEDLINE	ABB=ON	BETA-GLUCANS/CT OR GLUCANS/CT
L52	4	SEA	FILE=MEDLINE	ABB=ON	(L43 OR L44 OR L45) AND L51

=> s 146 or 152

L178 5 L46 OR L52

=> fil embase; d que 175; d que 169

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FILE COVERS 1974 TO 2 Feb 2006 (20060202/ED)

EMBASE has been reloaded. Enter HELP RLOAD for details.

This file contains CAS Registry Numbers for easy and accurate substance identification.

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L8
             1 SEA FILE=REGISTRY ABB=ON 37361-00-5
            2 SEA FILE=REGISTRY ABB=ON "B-D-GLUCAN, (1.FWDARW.3)-"/CN
L9
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L66
           15 SEA FILE=EMBASE ABB=ON GAULT R?/AU
L67
L68
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L75
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L68
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L69
=> fil drugu; d que 197;d que 1103
FILE 'DRUGU' ENTERED AT 13:05:48 ON 03 FEB 2006
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FILE LAST UPDATED: 31 JAN 2006
                                    <20060131/UP>
>>> DERWENT DRUG FILE (SUBSCRIBER) <<<
>>> FILE COVERS 1983 TO DATE <<<
>>> THESAURUS AVAILABLE IN /CT <<<
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L94
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L98
           80 SEA FILE=DRUGU ABB=ON GLUCAN-BETA-1,3/CT OR GLUCAN-BETA-1,3-D/
L99
               CT
            2 SEA FILE=DRUGU ABB=ON GLUCAN-BETA-1,6-D/CT
L100
             4 SEA FILE=DRUGU ABB=ON GLUCAN-BETA/CT
L101
             O SEA FILE=DRUGU ABB=ON (L94 OR L95 OR L96) AND (L98 OR L99 OR
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=> fil wpids; d que 1115; d que 1126

FILE 'WPIDS' ENTERED AT 13:05:49 ON 03 FEB 2006 COPYRIGHT (C) 2006 THE THOMSON CORPORATION

1 FEB 2006 FILE LAST UPDATED: <20060201/UP> MOST RECENT DERWENT UPDATE: 200608 <200608/DW> DERWENT WORLD PATENTS INDEX SUBSCRIBER FILE, COVERS 1963 TO DATE

>>> FOR A COPY OF THE DERWENT WORLD PATENTS INDEX STN USER GUIDE, PLEASE VISIT:

http://www.stn-international.de/training\_center/patents/stn\_guide.pdf <<<

- >>> FOR DETAILS OF THE PATENTS COVERED IN CURRENT UPDATES, SEE http://scientific.thomson.com/support/patents/coverage/latestupdates/
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http://scientific.thomson.com/support/products/dwpifv/

- >>> THE CPI AND EPI MANUAL CODES WILL BE REVISED FROM UPDATE 200601. PLEASE CHECK:
- http://scientific.thomson.com/support/patents/dwpiref/reftools/classification
- >>> PLEASE BE AWARE OF THE NEW IPC REFORM IN 2006, SEE

OR L125)

L112

- http://www.stn-international.de/stndatabases/details/ipc reform.html and
- http://scientific.thomson.com/media/scpdf/ipcrdwpi.pdf <<<

123 SEA FILE=WPIDS ABB=ON HUNTER K?/AU

L113	24	SEA	FILE=WPIDS	ABB=ON	GAULT R?/AU
L114	68	SEA	FILE=WPIDS	ABB=ON	JORDAN F?/AU
L115	2	SEA	FILE=WPIDS	ABB=ON	L112 AND L113 AND L114
L112	123	SEA	FILE=WPIDS	ABB=ON	HUNTER K?/AU
L113	24	SEA	FILE=WPIDS	ABB=ON	GAULT R?/AU
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L118	13730	SEA	FILE=WPIDS	ABB=ON	ADJUVANT#
L119	5852	SEA	FILE=WPIDS	ABB=ON	IMMUNOSTIMULA?
L120	436	SEA	FILE=WPIDS	ABB=ON	IMMUNOPOTENTIAT?
L121	483	SEA	FILE=WPIDS	ABB=ON	COSTIMULA? OR CO STIMULA?
L122	2049	SEA	FILE=WPIDS	ABB=ON	IMMUN#(W) (STIMULA? OR POTENTIAT? OR
		MODI	JLAT?)		
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L125					T(W) (CELL# OR LYMPHOCYTE#)
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	_				

(L117 OR L118 OR L119 OR L120 OR L121 OR L122 OR L123 OR L124

=> s 1115 or 1126

L179 2 L115 OR L126

=> fil biosis; d que 1140; d que 1143

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FILE COVERS 1969 TO DATE. CAS REGISTRY NUMBERS AND CHEMICAL NAMES (CNs) PRESENT FROM JANUARY 1969 TO DATE.

RECORDS LAST ADDED: 1 February 2006 (20060201/ED)

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L143	5	SEA :	FILE=BIOSIS ABB=ON (L137 OR L138 OR L139) AND (L141 OR
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=> s 1140 or 1143

L180 6 L140 OR L143

=> fil stnquide

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FILE CONTAINS CURRENT INFORMATION.
LAST RELOADED: Jan 27, 2006 (20060127/UP).

=> dup rem 1178,197,1177,1180,175,1179 FILE 'MEDLINE' ENTERED AT 13:06:33 ON 03 FEB 2006

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PROCESSING COMPLETED FOR L178
PROCESSING COMPLETED FOR L97
PROCESSING COMPLETED FOR L177
PROCESSING COMPLETED FOR L180
PROCESSING COMPLETED FOR L75
PROCESSING COMPLETED FOR L179

L181 11 DUP REM L178 L97 L177 L180 L75 L179 (9 DUPLICATES REMOVED)

ANSWERS '1-5' FROM FILE MEDLINE ANSWERS '6-8' FROM FILE CAPLUS ANSWERS '9-11' FROM FILE BIOSIS

=> d iall 1-5; d ibib ed abs hitind 6-8; d iall 9-11

L181 ANSWER 1 OF 11 MEDLINE on STN DUPLICATE 1

ACCESSION NUMBER: 2005158213 MEDLINE DOCUMENT NUMBER: PubMed ID: 15790516

TITLE: IFN-gamma primes macrophages for enhanced TNF-alpha

expression in response to stimulatory and non-stimulatory

amounts of microparticulate beta-glucan.

AUTHOR: Berner Mathew D; Sura Michael E; Alves Bryce N; Hunter

Kenneth W Jr

CORPORATE SOURCE: Department of Microbiology and Immunology, University of

Nevada School of Medicine, Applied Research Facility,

MS-199, Reno, NV 89557, USA.

SOURCE: Immunology letters, (2005 Apr 15) 98 (1) 15-22.

Electronic Publication: 2004 11-24. Journal code: 7910006. ISSN: 0165-2478.

PUB. COUNTRY: Netherlands

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200507

ENTRY DATE: Entered STN: 20050326

Last Updated on STN: 20050727 Entered Medline: 20050726

# ABSTRACT:

beta-(1-->3)-D-Glucan is an integral cell wall component of a variety of fungi, plants, and bacteria. Like the prototypic inflammatory mediator lipopolysaccharide (LPS), some beta-(1--> 3)-D-glucan-containing preparations have been shown to induce the production of proinflammatory cytokines by macrophages. In the present study, we have tested a new microparticulate form of beta-(1--> 3)-D-glucan (MG) from Saccharomyces cerevisiae for its ability to induce proinflammatory cytokine secretion in mouse peritoneal macrophages in vitro, and we have examined the effect of IFN-gamma. MG was rapidly phagocytized by peritoneal macrophages, and these MG-treated macrophages upregulated TNF-alpha, IL-6, and IL-1beta mRNAs and secreted these proinflammatory cytokines. IFN-gamma treatment alone did not induce unstimulated macrophages to produce TNF-alpha. However, a 4 h IFN-gamma pretreatment augmented TNF-alpha secretion by peritoneal macrophages subsequently treated with an optimally stimulatory dose of MG. IFN-gamma pretreatment for 2 h followed by thorough washing and a further 2 h incubation without IFN-gamma still resulted in enhanced TNF-alpha production in response

to MG, suggesting that IFN-gamma can prime macrophages for a subsequent proinflammatory response. Most interestingly, we found that IFN-gamma pretreatment of peritoneal macrophages enhanced the TNF-alpha response to amounts of MG that were poorly stimulatory or non-stimulatory in the absence of IFN-gamma priming. These data suggest that a synergy between IFN-gamma and beta-glucan may have evolved to lower the threshold of sensitivity of the innate immune response to fungal pathogens.

Check Tags: Female CONTROLLED TERM:

Animals

Cytokines: BI, biosynthesis Cytokines: GE, genetics Cytokines: SE, secretion

Gene Expression Regulation: PH, physiology

\*Interferon Type II: ME, metabolism Lipopolysaccharides: ME, metabolism \*Macrophages, Peritoneal: ME, metabolism Macrophages, Peritoneal: SE, secretion

Mice

Mice, Inbred BALB C

Phagocytosis: PH, physiology RNA, Messenger: ME, metabolism Research Support, Non-U.S. Gov't

Tumor Necrosis Factor-alpha: GE, genetics \*Tumor Necrosis Factor-alpha: ME, metabolism

\*beta-Glucans: ME, metabolism

CAS REGISTRY NO.:

82115-62-6 (Interferon Type II)

CHEMICAL NAME: 0 (Cytokines); 0 (Lipopolysaccharides); 0 (RNA, Messenger);

0 (Tumor Necrosis Factor-alpha); 0 (beta-1,3-D-glucan); 0

(beta-Glucans)

L181 ANSWER 2 OF 11 MEDLINE on STN DUPLICATE 3

ACCESSION NUMBER: 2004237405 MEDLINE DOCUMENT NUMBER: PubMed ID: 15134902

Microparticulate beta-glucan upregulates the expression of TITLE:

B7.1, B7.2, B7-H1, but not B7-DC on cultured murine

peritoneal macrophages.

Hunter Kenneth W Jr; DuPre' Sally; Redelman Doug AUTHOR:

Department of Microbiology and Immunology, University of CORPORATE SOURCE:

Nevada School of Medicine, Reno, NV 89557, USA...

khunter@unr.edu

CONTRACT NUMBER: P20 RR16464 (NCRR)

SOURCE:

(2004 Apr 30) 93 (1) 71-8. Immunology letters,

Journal code: 7910006. ISSN: 0165/2478.

PUB. COUNTRY: Netherlands

(YOURNAL ARTICLE) DOCUMENT TYPE: Journal; Article;

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200412

Entered STN: 20040512 ENTRY DATE:

> Last Updated on STN: 20041222 Entered Medline: 20041221

# ABSTRACT:

Beta-1,3-(D)-glucan from a variety of biological sources has been shown to enhance both humoral and cellular immune responses to a variety of antigens, infectious agents, and tumors. Nevertheless, its mode of action has not been fully defined. We sought to determine whether a 1-2 microm diameter microparticulate form of beta-glucan (MG) from the yeast Saccharomyces cerevisiae could regulate expression of B7 family glycoproteins on resident peritoneal macrophages from BALB/c mice. We discovered that MG uregulated B7.2 mRNA expression and enhanced the surface membrane expression of B7.2

qlycoprotein. Although B7.1 mRNA was not upregulated above constitutive levels, MG treatment enhanced B7.1 glycoprotein expression on the macrophages, albeit to a lesser extent than B7.2. At the same time, the gene and cell surface expression of B7-H1, a putative negative regulator of T cell activity, was also upregulated by MG. The expression of B7-DC, another B7 family molecule with negative regulatory activity, was not affected by incubation with This study has demonstrated that a microparticulate form of beta-glucan can enhance B7 co-stimulatory molecule expression on macrophages, thereby enabling these antigen-presenting cells to deliver the second signal to T-lymphocytes that express CD28. In addition, because MG also induces the expression of B7-H1, it may enable macrophages to provide a concomitant downregulatory signal to T-lymphocytes expressing PD-1 or related receptors.

CONTROLLED TERM: Check Tags: Female

Animals

Antigens, CD: GE, genetics \*Antigens, CD: IM, immunology Antigens, CD: ME, metabolism Antigens, CD80: GE, genetics \*Antigens, CD80: IM, immunology Antigens, CD80: ME, metabolism Blood Proteins: GE, genetics \*Blood Proteins: IM, immunology Blood Proteins: ME, metabolism

Gene Expression Regulation: PH, physiology \*Macrophages, Peritoneal: IM, immunology Membrane Glycoproteins: GE, genetics \*Membrane Glycoproteins: IM, immunology Membrane Glycoproteins: ME, metabolism

Mice

Peptides: GE, genetics \*Peptides: IM, immunology Peptides: ME, metabolism RNA, Messenger: ME, metabolism Research Support, Non-U.S. Gov't Research Support, U.S. Gov't, P.H.S.

Reverse Transcriptase Polymerase Chain Reaction

Up-Regulation

\*beta-Glucans: ME, metabolism

CHEMICAL NAME:

0 (Antigens, CD); 0 (Antigens, CD80); 0 (B7-DC antigen); 0 (Blood Proteins); 0 (CD86 antigen); 0 (Membrane Glycoproteins); 0 (PDCD1LG1 protein, human); 0 (Peptides); 0 (RNA, Messenger); 0 (beta-1,3-D-glucan); 0 (beta-Glucans)

MEDLINE on STN **DUPLICATE 4** L181 ANSWER 3 OF 11

ACCESSION NUMBER: 2003021273 MEDLINE DOCUMENT NUMBER: PubMed ID: 12526860

TITLE: Synthesis of cetyl myristoleate and evaluation of its

therapeutic efficacy in a murine model of collagen-induced

arthritis.

Hunter Kenneth W Jr; Gault Ruth A; **AUTHOR:** 

Stehouwer Jeffrey S; Tam-Chang Suk-Wah

CORPORATE SOURCE: Department of Microbiology, University of Nevada School of

Medicine, Reno, NV 89557, USA.. khunter@unr.edu

Pharmacological research : official journal of the Italian SOURCE:

Pharmacological Society, (2003 Jan) 47 (1) 43-7. Journal code: 8907422. ISSN: 1043-6618.

PUB. COUNTRY: England: United Kingdom

Journal; Article; (JOURNAL ARTICLE) DOCUMENT TYPE:

LANGUAGE: English

Priority Journals FILE SEGMENT:

Jones 10/630143

Page 9

ENTRY MONTH:

200310

ENTRY DATE:

Entered STN: 20030116

Last Updated on STN: 20031008 Entered Medline: 20031006

## ABSTRACT:

Cetyl myristoleate (CM) was reported by Diehl and May [J Pharm Sci 83 (1994) 296] to block inflammation and prevent adjuvant-induced arthritis in rats. To verify this earlier work, we have synthesized pure CM and tested its anti-arthritic properties in a collagen-induced arthritis model in DBA/1LacJ mice. Multiple intraperitoneal injections of CM in 450 and 900 mg kg(-1) doses resulted in a significantly lower incidence of disease and caused a modest but significant diminution in clinical signs in those mice that developed arthritis. CM administered in daily oral doses of 20 mg kg(-1) also reduced the incidence of arthritis and caused a small reduction in the clinical signs in mice that developed arthritis. Although the protective effect of CM in collagen-induced arthritis observed in the present study was less dramatic than that reported earlier, our results confirm the anti-arthritic properties of pure CM.

CONTROLLED TERM:

Check Tags: Comparative Study; Female

Animals

\*Arthritis, Experimental: DT, drug therapy Arthritis, Experimental: PA, pathology

\*Disease Models, Animal

Drug Evaluation, Preclinical: MT, methods

Mice

Mice, Inbred DBA

Research Support, Non-U.S. Gov't \*Waxes: CS, chemical synthesis \*Waxes: TU, therapeutic use

CAS REGISTRY NO.:

64660-84-0 (cetyl myristoleate)

CHEMICAL NAME: 0 (Waxes)

L181 ANSWER 4 OF 11

MEDLINE on STN

DUPLICATE 6

ACCESSION NUMBER: 2002498220 MEDLINE PubMed ID: 12358685

DOCUMENT NUMBER:

TITLE: Preparation of microparticulate beta-glucan from

Saccharomyces cerevisiae for use in immune potentiation.

Hunter K W Jr; Gault R A; Berner M D AUTHOR:

CORPORATE SOURCE:

Department of Microbiology, University of Nevada School of

Medicine, Reno, NV 89557, USA. . Khunter@unr.edu

SOURCE:

Letters in applied microbiologý, (2002) 35 (4) 267-71.

Journal code: 8510094. ISSN: /0266-8254.

PUB. COUNTRY:

England: United Kingdom

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

200212

ENTRY DATE:

Entered STN: 20021003

Last Updated on STN: 20021217 Entered Medline: 20021204

#### ABSTRACT:

AIMS: To develop a method for the preparation of an immunologically active, homogeneous, nonaggregated, microparticulate beta-glucan-containing material from the budding yeast Saccharomyces cerevisiae. METHODS AND RESULTS: Using a combination of sonication and spray-drying, a homogeneous preparation of 1-2-mu diameter beta-glucan-containing particles was made from alkali- and acid-insoluble yeast cell wall material. This microparticulate beta-glucan remained in suspension longer and, following oral administration at 0.1 mg kg(-1) for 14 d, enhanced phagocytosis of mouse peritoneal macrophages significantly better than did aggregated beta-glucan particles. CONCLUSIONS: A

new sonication and spray-drying method can be employed to overcome the problem of aggregation of beta-glucan microparticles in aqueous media. SIGNIFICANCE AND IMPACT OF THE STUDY: A microparticulate form of beta-glucan that remains in suspension longer for pharmaceutical applications and has superior immune potentiation characteristics has been developed.

CONTROLLED TERM: Check Tags: In Vitro

\*Adjuvants, Immunologic: IP, isolation & purification

Adjuvants, Immunologic: PD, pharmacology

Animals

Glucans: IM, immunology

\*Glucans: IP, isolation & purification

Glucans: PD, pharmacology
Macrophages: DE, drug effects
Macrophages: IM, immunology

Mice

Mice, Inbred BALB C

Phagocytosis: DE, drug effects

Reagent Kits, Diagnostic

Research Support, Non-U.S. Gov't

\*Saccharomyces cerevisiae: CH, chemistry

Sonication

CHEMICAL NAME: 0 (Adjuvants, Immunologic); 0 (Glucans); 0 (Reagent Kits,

Diagnostic)

L181 ANSWER 5 OF 11 MEDLINE on STN ACCESSION NUMBER: 97253676 MEDLINE DOCUMENT NUMBER: PubMed ID: 9099058

TITLE: The impact of non-endotoxin LAL-reactive materials on

Limulus amebocyte lysate analyses.

AUTHOR: Cooper J F; Weary M E; Jordan F T

CORPORATE SOURCE: Charles River Endosafe, Charleston, South Carolina, USA.

SOURCE: PDA journal of pharmaceutical science and technology / PDA,

(1997 Jan-Feb) 51 (1) 2-6. Ref: 32 Journal code: 9439538. ISSN: 1079-7440.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

General Review; (REVIEW)

(REVIEW, TUTORIAL)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199707

ENTRY DATE: Entered STN: 19970721

Last Updated on STN: 19970721 Entered Medline: 19970703

## ABSTRACT:

Limulus amebocyte lysate (LAL) is activated by bacterial endotoxins and certain glucans. (beta-D-glucan, LAL-RM). The potential for conflicting inter-laboratory results for LAL tests exists because commercial LAL reagents are highly variable in response to LAL-reactive glucans. The nature of beta-D-glucan activation of LAL and means for rendering LAL non-responsive to glucan are reviewed to provide a background for resolving conflicting data. Kinetic LAL methods are particularly useful for screening materials potentially contaminated with glucan. The presence of beta-D-glucan in parenterals is uncommon and is likely limited to products exposed to microbial or cellulosic materials. A scheme is suggested for identifying LAL-reactive glucans and for LAL release-testing without glucan interference.

CONTROLLED TERM: Glucans: AN, analysis

Indicators and Reagents \*Limulus Test: MT, methods

CHEMICAL NAME: 0 (Glucans); 0 (Indicators and Reagents)

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L181 ANSWER 6 OF 11 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 2
ACCESSION NUMBER:
                               2004:120668 CAPLUS
DOCUMENT NUMBER:
                               140:162361
TITLE:
                               compositions containing microparticulate
                               \beta-1,3(6) - glucan and conjugates for use
                               as vaccine adjuvants
INVENTOR(S):
                               Hunter, Kenneth W.; Jordan, Frank M.
                               ; Gault, Ruth A.
PATENT ASSIGNEE(S):
                               Immusonic, Inc., USA
SOURCE:
                               PCT Int. Appl., 61 pp.
                               CODEN: PIXXD2
DOCUMENT TYPE:
                               Patent
                               English
LANGUAGE:
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
                           KIND
                                                     APPLICATION NO.
      PATENT NO.
                                       DATE
                               ----
                                        -----
                                                       -----
                                                                                    -----
      WO 2004012657
                               A2
                                        20040212
                                                     WO 2003-US23741
                                                                                    20030730
                               A3
      WO 2004012657
                                       20040708
           W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
                CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
                GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
          PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW

RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR,
                \mathtt{BF},\ \mathtt{BJ},\ \mathtt{CF},\ \mathtt{CG},\ \mathtt{CI},\ \mathtt{CM},\ \mathtt{GA},\ \mathtt{GN},\ \mathtt{GQ},\ \mathtt{GW},\ \mathtt{ML},\ \mathtt{MR},\ \mathtt{NE},\ \mathtt{SN},\ \mathtt{TD},\ \mathtt{TG}
PRIORITY APPLN. INFO.:
                                                      US 2002-400377P
                                                                              P 20020801
ED
      Entered STN: 13 Feb 2004
ΔR
      A microparticulate beta-glucan is used as a vaccine adjuvant for animals
      and humans, binding to qlucan receptors on a variety of phagocytic cells
      to enhance their immunol. functions. The particles contain about 1-10%
      partially deacetylated N-acetylglucosamine and are predominantly 0.3-3 \mu in diameter, preferably 1 - 2 \mu in diameter, to cause the expression of
      co-stimulatory mols. on antigen presenting cells (APC's). The
      microparticle upregulates the expression of the co-stimulatory mol.
      B7 based upon such microparticles containing beta- (1,3) and beta
      (1,6) glucan.
IC
      ICM A61K
      15-2 (Immunochemistry)
CC
      Section cross-reference(s): 9, 63
ST
      beta glucan microparticulate vaccine adjuvant
      B7 antigen presenting cell
IT
      Vaccines
          (AIDS; compns. containing microparticulate \beta-1,3- or \beta-1,6-
         glucan and conjugates for use as vaccine adjuvants)
TT
      Antigens
      RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
      (Biological study); USES (Uses)
          (B7-3; compns. containing microparticulate \beta-1,3- or
         \beta-1,6- glucan and conjugates for use as vaccine
         adjuvants)
      Hematopoiesis
IT
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(T-cell lymphopoiesis; compns. containing microparticulate \beta-1,3- or
        \beta-1,6- glucan and conjugates for use as vaccine
        adjuvants)
ΙT
     Macrophage
         (activation; compns. containing microparticulate \beta-1,3- or \beta-1,6-
        glucan and conjugates for use as vaccine adjuvants)
IT
     Immunostimulants
         (adjuvants; compns. containing microparticulate \beta-1,3- or
        \beta-1,6- glucan and conjugates for use as vaccine
        adjuvants)
IT
     Vaccines
         (antimalarial; compns. containing microparticulate \beta-1,3- or
        \beta-1,6- glucan and conjugates for use as vaccine
        adjuvants)
ΙT
     Biochemical compounds
         (co-stimulatory; compns. containing microparticulate \beta-1,3- or
        \beta-1,6- glucan and conjugates for use as vaccine
        adjuvants)
IT
     Amino group
     Animal
     Antigen-presenting cell
     Dehydration, physiological
     Drying
     Freeze drying
     Grinding (size reduction)
     Human
     Infection
     Macrophage
     Saccharomyces
     Sonication
     Spraying
     Spraying apparatus
       T cell (lymphocyte)
       Vaccines
     Veterinary medicine
         (compns. containing microparticulate \beta-1,3- or \beta-1,6-
        glucan and conjugates for use as vaccine adjuvants)
IT
     Antigens
     CD80 (antigen)
     CD86 (antigen)
     Lipids, biological studies
     Oligosaccharides, biological studies
     Polysaccharides, biological studies
     Proteins
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
         (compns. containing microparticulate \beta-1,3- or \beta-1,6-
        glucan and conjugates for use as vaccine adjuvants)
IT
     Lymphocyte
         (effector cell; compns. containing microparticulate \beta-1,3- or
        \beta-1,6- glucan and conjugates for use as vaccine
        adjuvants)
IT
     Organelle
        (globule; compns. containing microparticulate \beta-1,3- or \beta-1,6-
        glucan and conjugates for use as vaccine adjuvants)
     Drug delivery systems
IT
        (liqs.; compns. containing microparticulate \beta-1,3- or \beta-1,6-
        glucan and conjugates for use as vaccine adjuvants)
     Cell activation
TΤ
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Peritoneum
        (macrophage; compns. containing microparticulate \beta-1,3- or \beta-1,6-
        glucan and conjugates for use as vaccine adjuvants)
IT
     Drug delivery systems
        (microparticles; compns. containing microparticulate \beta-1,3- or
        \beta-1,6- glucan and conjugates for use as vaccine
        adjuvants)
IT
     Drug delivery systems
        (particles; compns. containing microparticulate \beta-1,3- or \beta-1,6-
        glucan and conjugates for use as vaccine adjuvants)
IT
     Macrophage
        (peritoneal; compns. containing microparticulate \beta-1,3- or \beta-1,6-
        glucan and conjugates for use as vaccine adjuvants)
IT
     Hydration, physiological
        (rehydration; compns. containing microparticulate \beta-1,3- or
        \beta-1,6- glucan and conjugates for use as vaccine
        adjuvants)
IT
     Infection
        (schistosomiasis; compns. containing microparticulate β-1,3- or
        \beta-1,6- glucan and conjugates for use as vaccine
        adjuvants)
IT
     Coating process
        (spray; compns. containing microparticulate \beta-1,3- or \beta-1,6-
        glucan and conjugates for use as vaccine adjuvants)
IT
     Anti-AIDS agents
     Antimalarials
        (vaccines; compns. containing microparticulate \beta-1,3- or \beta-1,6-
        glucan and conjugates for use as vaccine adjuvants)
     50-99-7, Glucose, biological studies 1398-61-4, Chitin
IT
     2280-44-6D, D-Glucopyranose, \beta-1,3- and \beta-1,6- derivs.
     7512-17-6D, N-Acetylglucosamine, partially deacetylated derivs.
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (compns. containing microparticulate \beta-1,3- or \beta-1,6-
        glucan and conjugates for use as vaccine adjuvants)
L181 ANSWER 7 OF 11 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 5
ACCESSION NUMBER:
                         2002:813870 CAPLUS
DOCUMENT NUMBER:
                         137:299970
TITLE:
                         The use of beta-1,3-glucan-containing
                         compositions to potentiate immune responses by
                         upregulating the expression of costimulatory molecules
INVENTOR (S):
                         Hunter, Kenneth W.; Gault, Ruth A.
                         ; Jordan, Frank M.
PATENT ASSIGNEE(S):
                         Immusonic, Inc., USA
SOURCE:
                         PCT Int. Appl., 37 pp.
                         CODEN: PIXXD2
DOCUMENT TYPE:
                         Patent
LANGUAGE:
                         English
FAMILY ACC. NUM. COUNT: 3
PATENT INFORMATION:
     PATENT NO.
                        KIND
                                DATE APPLICATION NO. DATE
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                                _____
                                             ______
     _____
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    WO 2002083061 A2 20021024
WO 2002083061 A3 20030103
                                            WO 2001-US43711
                                                                    20011106
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
             CO, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM,
             HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS,
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LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO,

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RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ,
             VN, YU, ZA, ZW
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
             DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
             BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
     US 6476003
                                20021105
                                             US 2000-707583
                          B1
                                                                    20001106
PRIORITY APPLN. INFO.:
                                             US 2000-707436
                                                                 A 20001106
                                             US 2000-707437
                                                                A 20001106
                                                                 A 20001106
                                             US 2000-707582
                                             US 2000-707583
                                                                A 20001106
ED
     Entered STN: 25 Oct 2002
     An improved method and immunopharmacol. composition for upregulating the
AB
     expression of the co-stimulatory mol. B7 is provided. A
     β-1,3-glucan-containing composition is provided that can upregulate the cell
     surface expression of B7 mols. on antigen presenting cells like
     macrophages, thereby allowing these antigen presenting cells to more
     effectively initiate adaptive immune responses to foreign antigens like
     pathogenic microorganisms and tumors.
IC
     ICM A61K
     63-6 (Pharmaceuticals)
CC
     Section cross-reference(s): 1
ST
     glucan B7 antigen upregulation immunostimulant tumor
     microorganism
     Antibodies and Immunoglobulins
IT
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (IqM; use of \beta-1,3- glucan-containing compns. to potentiate
        immune responses by upregulating expression of costimulatory mols.)
IT
     Respiration, animal
        (burst; use of \beta-1,3- glucan-containing compns. to potentiate
        immune responses by upregulating expression of costimulatory mols.)
IT
     Drug delivery systems
        (capsules; use of \beta-1,3- glucan-containing compns. to
        potentiate immune responses by upregulating expression of costimulatory
        mols.)
IT
     T cell (lymphocyte)
        (effector; use of \beta-1,3- glucan-containing compns. to
        potentiate immune responses by upregulating expression of costimulatory
        mols.)
IT
     Drug delivery systems
        (globules; use of \beta-1,3- glucan-containing compns. to
        potentiate immune responses by upregulating expression of costimulatory
        mols.)
IT
    Drug delivery systems
        (liqs.; use of \beta-1,3- glucan-containing compns. to potentiate
        immune responses by upregulating expression of costimulatory mols.)
IT
     Cell differentiation
        (lymphocyte; use of \beta-1,3- glucan-containing compns. to
        potentiate immune responses by upregulating expression of costimulatory
        mols.)
IT
        (macrophage; use of \beta-1,3- glucan-containing compns. to
        potentiate immune responses by upregulating expression of costimulatory
        mols.)
IT
    Macrophage
        (peritoneal; use of \beta-1,3- glucan-containing compns. to
        potentiate immune responses by upregulating expression of costimulatory
        mols.)
IT
    Drug delivery systems
        (powders; use of \beta-1,3- glucan-containing compns. to
        potentiate immune responses by upregulating expression of costimulatory
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mols.)
IT
     Drug delivery systems
        (tablets; use of \beta-1,3- glucan-containing compns. to
        potentiate immune responses by upregulating expression of costimulatory
        mols.)
TΤ
     CD80 (antigen)
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (upregulation of expression of; use of \beta-1,3- glucan
        -containing compns. to potentiate immune responses by upregulating
        expression of costimulatory mols.)
TΤ
     B cell (lymphocyte)
     Dendritic cell
       Immunostimulants
     Macrophage
     Particle size
     Phagocytosis
     Sonication
        (use of \beta-1,3- glucan-containing compns. to potentiate immune
        responses by upregulating expression of costimulatory mols.)
TT
     Microorganism
     Neoplasm
        (use of \beta-1,3- glucan-containing compns. to potentiate immune
        responses to microorganisms and tumor by upregulating expression of
        costimulatory mols.)
IT
     9051-97-2 37361-00-5, β-1,6- Glucan
     RL: DMA (Drug mechanism of action); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (use of \beta-1,3- glucan-containing compns. to potentiate immune
        responses by upregulating expression of costimulatory mols.)
L181 ANSWER 8 OF 11 CAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER:
                         2004:533968 CAPLUS
DOCUMENT NUMBER:
                         141:70241
TITLE:
                         Compositions comprising \beta(1,3) - glucans
                         and \beta(1,6) - glucans for use as vaccine
                         adjuvant and methods of manufacturing \beta-
                         glucans
INVENTOR(S):
                         Hunter, Kenneth W.; Gault, Ruth A.
                         ; Jordan, Frank M.
PATENT ASSIGNEE(S):
                         USA
                         U.S. Pat. Appl. Publ., 26 pp., Cont.-in-part of U\S.
SOURCE:
                         Ser. No. 707,582.
                         CODEN: USXXCO
DOCUMENT TYPE:
                         Patent
LANGUAGE:
                         English
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
     PATENT NO.
                         KIND
                                DATE
                                             APPLICATION NO
                                                                     DATE
     ______
                         ____
                                 -----
                                             -----
    US 2004127458
                          A1
                                20040701
                                             US 2003-630143
                                                                     20030730 <--
PRIORITY APPLN. INFO.:
                                             US 2000-707582
                                                                  A2 20001106
                                             US 2002-400377P
                                                                  P 20020801
```

ED Entered STN: 02 Jul 2004

AB A microparticulate beta-glucan is used as a vaccine adjuvant for animals and humans, binding to glucan receptors on a variety of phagocytic cells to enhance their immunol. functions. The particles contain about 1-10% partially deacetylated N-acetylglucosamine and are predominantly 0.3-3  $\mu$  in diameter, preferably 1-2  $\mu$  in diameter, to cause the expression of co-stimulatory mols. on antigen presenting cells (APC's). The

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microparticle upregulates the expression of the co-stimulatory mol.
     B7, based upon such microparticles containing beta-(1,3) and beta(1,6)
     glucan.
     ICM A61K031-715
INCL 514054000
     15-2 (Immunochemistry)
CC
     Section cross-reference(s): 63
     beta glucan vaccine adjuvant antigen conjugate
ST
     microparticle phagocyte APC
TT
     Antigens
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
         (B7.3; compns. comprising \beta(1,3) - glucans and
        \beta(1,6) - glucans for use as vaccine adjuvant and
        methods of manufacturing \beta- glucans)
IT
     Hematopoiesis
        (T-cell lymphopoiesis; compns. comprising \beta(1,3) - glucans
        and \beta(1,6) - glucans for use as vaccine adjuvant
        and methods of manufacturing \beta- glucans)
IT
     Macrophage
         (activation; compns. comprising \beta(1,3) - glucans and
        \beta(1,6) - glucans for use as vaccine adjuvant and
        methods of manufacturing \beta- glucans)
IT
     Immunostimulants
         (adjuvants; compns. comprising \beta(1,3) - glucans
        and \beta(1,6) - glucans for use as vaccine adjuvant
        and methods of manufacturing β- glucans)
IT
     Molecules
         (co-stimulatory; compns. comprising \beta(1,3) - glucans and
        \beta(1,6) - glucans for use as vaccine adjuvant and
        methods of manufacturing \beta- glucans)
IT
     Amino group
     Animal
     Antigen-presenting cell
     Grinding (size reduction)
     Human
     Microparticles
     Phagocyte
     Signal transduction, biological
     Sonication
     Spraying apparatus
       T cell (lymphocyte)
       Vaccines
         (compns. comprising \beta(1,3) - glucans and \beta(1,6) -
        glucans for use as vaccine adjuvant and methods of
        manufacturing \beta- glucans)
IT
     CD80 (antigen)
     CD86 (antigen)
     Gelatins, biological studies
     Polysaccharides, biological studies
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (compns. comprising \beta(1,3) - glucans and \beta(1,6) -
        glucans for use as vaccine adjuvant and methods of
        manufacturing \beta- glucans)
IT
     Antigens
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (conjugates; compns. comprising \beta(1,3) - glucans and
        \beta(1,6) - glucans for use as vaccine adjuvant and
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methods of manufacturing \beta- glucans)
     T cell (lymphocyte)
TT
        (effector cell; compns. comprising \beta(1,3) - glucans and
        \beta(1,6) - glucans for use as vaccine adjuvant and
        methods of manufacturing \beta- glucans)
ΤТ
     Receptors
     RL: BSU (Biological study, unclassified); BIOL (Biological study)
        (glucan; compns. comprising \beta(1,3) - glucans and
        \beta(1,6) - glucans for use as vaccine adjuvant and
        methods of manufacturing \beta- glucans)
     Drug delivery systems
TT
        (microparticles; compns. comprising \beta(1,3) - glucans and
        \beta(1,6) - glucans for use as vaccine adjuvant and
        methods of manufacturing \beta- glucans)
     Hydration, physiological
IT
        (rehydration, without reaggregation; compns. comprising \beta(1,3)-
        glucans and \beta(1,6) - glucans for use as vaccine
        adjuvant and methods of manufacturing β- glucans)
     Particles
IT
        (small; compns. comprising \beta(1,3) - glucans and
        \beta(1,6) - glucans for use as vaccine adjuvant and
        methods of manufacturing \beta- glucans)
IT
     Drying
        (without reaggregation; compns. comprising \beta(1,3) - glucans
        and \beta(1,6) - glucans for use as vaccine adjuvant
        and methods of manufacturing \beta- glucans)
     1398-61-4, Chitin 7512-17-6D, N-Acetylglucosamine,
IT
     deacylated derivs. 9051-97-2D, analogs 37361-00-5D,
     \beta(1,6) - Glucan, analogs
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (compns. comprising \beta(1,3) - glucans and \beta(1,6) -
        glucans for use as vaccine adjuvant and methods of
        manufacturing \beta- glucans)
```

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ACCESSION NUMBER:
                    2003:32049 BIOSIS
DOCUMENT NUMBER:
                    PREV200300032049
TITLE:
                    Method for preparing small particle size glucan in a dry
AUTHOR (S):
                    Jordan, Frank M. [Inventor, Reprint Author];
                    Gault, Ruth A. [Inventor]; Hunter, Kenneth
                    W. [Inventor]
CORPORATE SOURCE:
                    Reno, NV, USA
                    ASSIGNEE: Immusonic, Inc., Carson City, NV, USA
PATENT INFORMATION: US 6476003 20021105
SOURCE:
                    Official Gazette of the United States Patent and Trademark
                    Office Patents, (Nov 5 2002) Vol. 1264, No. 1.
                    http://www.uspto.gov/web/menu/patdata.html. e-file.
                    ISSN: 0098-1133 (ISSN print).
DOCUMENT TYPE:
                    Patent
LANGUAGE:
                    English
ENTRY DATE:
                    Entered STN: 8 Jan 2003
                    Last Updated on STN: 8 Jan 2003
```

ABSTRACT: An improved method for purifying glucan to small particle size glucan and drying the glucan to a solid such that the glucan may be re-hydrated and maintain substantially maintain a particle size of one micron or less so that

L181 ANSWER 9 OF 11 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN

it may be used in nutritional, pharmaceutical and pharmacological compositions where a dry material is desired such that a greater immunological benefit may be obtained.

NAT. PATENT. CLASSIF.:514054000

Biochemistry studies - General CONCEPT CODE: 10060

> Biochemistry studies - Carbohydrates 10068

INDEX TERMS: Major Concepts

Biochemistry and Molecular Biophysics; Methods and

Techniques

Chemicals & Biochemicals INDEX TERMS:

glucan: purification

Methods & Equipment INDEX TERMS:

small particle size glucan preparation: laboratory

techniques

REGISTRY NUMBER: 9012-72-0 (glucan)

L181 ANSWER 10 OF 11 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on

ACCESSION NUMBER: 2002:348087 BIOSIS PREV200200348087 DOCUMENT NUMBER:

Methods and compositions for the detection of bacterial TITLE:

endotoxins.

Jordan, Foster T. [Inventor, Reprint author]; AUTHOR(S):

Chiang, Hui-Ti [Inventor]; Cooper, James F. [Inventor];

Wainwright, Norman R. [Inventor]

Hollywood, SC, USA CORPORATE SOURCE:

ASSIGNEE: Charles River Laboratories

PATENT INFORMATION: US 6391570 20020521

Official Gazette of the United States Patent and Trademark SOURCE:

Office Patents, (May 21, 2002) Vol. 1258, No. 3. http://www.uspto.gov/web/menu/patdata.html. e-file.

CODEN: OGUPE7. ISSN: 0098-1133.

DOCUMENT TYPE:

Patent LANGUAGE: English

ENTRY DATE: Entered STN: 19 Jun 2002

Last Updated on STN: 19 Jun 2002

ABSTRACT: The invention provides methods and compositions for the detection and/or quantification of bacterial endotoxins. In particular, provided herein is an inexpensive and reproducible method for producing an improved amebocyte lysate preparation having reduced Factor G activity. Provided also is an endotoxin-specific amebocyte lysate preparation produced by such a method. addition, the invention provides methods and compositions akin for enhancing the sensitivity to endotoxins of amebocyte lysate preparations having reducing Factor G activity. In particular, the sensitivity of such amebocyte lysate preparations to endotoxins can be enhanced by the addition of exogenous (1fwdarw3) beta-D-glucan.

NAT. PATENT. CLASSIF.:435732000

CONCEPT CODE: 12504

Pathology - Diagnostic Pathology - Therapy 12512 Pharmacology - General 22002

Physiology and biochemistry of bacteria 31000

INDEX TERMS: Major Concepts

Methods and Techniques; Pharmacology

INDEX TERMS: Chemicals & Biochemicals

Factor G; bacterial endotoxin detection compositions:

diagnostic-drug; bacterial endotoxins; beta-D-

glucan

INDEX TERMS: Methods & Equipment

bacterial endotoxin detection: detection method

ORGANISM: Classifier Jones 10/630143

Bacteria 05000

Super Taxa

Microorganisms
Organism Name
bacteria
Taxa Notes

Bacteria, Eubacteria, Microorganisms

REGISTRY NUMBER:

23297-71-4 (Factor G) 9041-22-9 (beta-D-glucan)

L181 ANSWER 11 OF 11 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on

STN

ACCESSION NUMBER: 2001:457213 BIOSIS DOCUMENT NUMBER: PREV200100457213

TITLE: Methods and compositions for the detection of bacterial

endotoxins.

AUTHOR(S): Jordan, Foster T. [Inventor, Reprint author];

Chiang, Hui-Ti [Inventor]; Cooper, James F. [Inventor];

Wainwright, Norman R. [Inventor]

CORPORATE SOURCE: Hollywood, SC, USA

ASSIGNEE: Charles River Laboratories, Wilmington, MA, USA

PATENT INFORMATION: US 6270982 20010807

SOURCE:

Official Gazette of the United States Patent and Trademark Office Patents, (Aug. 7, 2001) Vol. 1249, No. 1. e-file.

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Patent English

LANGUAGE: ENTRY DATE:

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ABSTRACT: The invention provides methods and compositions for the detection and/or quantification of bacterial endotoxins. In particular, provided herein is an inexpensive and reproducible method for producing an improved amebocyte lysate preparation having reduced Factor G activity. Provided also is an endotoxin-specific amebocyte lysate preparation produced by such a method. In addition, the invention provides methods and compositions for enhancing the sensitivity to endotoxins of amebocyte lysate preparations having reducing Factor G activity. In particular, the sensitivity of such amebocyte lysate preparations to endotoxins can be enhanced by the addition of exogenous (1fwdarw3) beta-D-glucan.

NAT. PATENT. CLASSIF.:435732000

CONCEPT CODE: General biology - Miscellaneous 00532

INDEX TERMS: Major Concepts

Biochemistry and Molecular Biophysics; Methods and

Techniques

INDEX TERMS: Chemicals & Biochemicals

amebocyte lysate preparation: reduced Factor G activity;

bacterial endotoxins; exogenous (1-FAR3) beta

-D-glucan.

INDEX TERMS: Methods & Equipment

bacterial endotoxin detection: detection method; bacterial endotoxin quantification: quantification

method

=> fil capl; d que 128; d que 131; d que 142 FILE 'CAPLUS' ENTERED AT 13:09:14 ON 03 FEB 2006 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2006 AMERICAN CHEMICAL SOCIETY (ACS)

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BAC - Biological activity
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=> fil medl; d que 156; d que 157; d que 165

FILE 'MEDLINE' ENTERED AT 13:09:16 ON 03 FEB 2006

FILE LAST UPDATED: 2 FEB 2006 (20060202/UP). FILE COVERS 1950 TO DATE.

On December 11, 2005, the 2006 MeSH terms were loaded.

The MEDLINE reload for 2006 will soon be available. For details on the 2005 reload, enter HELP RLOAD at an arrow promt (=>). See also:

http://www.nlm.nih.gov/mesh/

http://www.nlm.nih.gov/pubs/techbull/nd04/nd04\_mesh.html

http://www.nlm.nih.gov/pubs/techbull/nd05/nd05\_med\_data\_changes.html

http://www.nlm.nih.gov/pubs/techbull/nd05/nd05\_2006\_MeSH.html

OLDMEDLINE is covered back to 1950.

MEDLINE thesauri in the /CN, /CT, and /MN fields incorporate the

MeSH 2006 vocabulary.

This file contains CAS Registry Numbers for easy and accurate

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18 (L56 OR L57 OR L65) NOT (L178

=> fil embase; d que 186; d que 193

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Searched by Barb O'Bryen, STIC 2-2518

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MOST RECENT DERWENT UPDATE: 200608 <200608/DW>
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L119	5852	SEA	FILE=WPIDS	ABB=ON	IMMUNOSTIMULA?
L120	436	SEA	FILE=WPIDS	ABB=ON	IMMUNOPOTENTIAT?
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FILE COVERS 1969 TO DATE. CAS REGISTRY NUMBERS AND CHEMICAL NAMES (CNs) PRESENT FROM JANUARY 1969 TO DATE.

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                  970 SEA FILE=BIOSIS ABB=ON (L8 OR L9)
             5632 SEA FILE=BIOSIS ABB=ON GLUCAN#(3A)BETA
271754 SEA FILE=BIOSIS ABB=ON T(W) (CELL# OR LYMPHOCYTE#)
18096 SEA FILE=BIOSIS ABB=ON IMMUNOSTIMULA?
18802 SEA FILE=BIOSIS ABB=ON IMMUNOMODULAT?
2 SEA FILE=BIOSIS ABB=ON (L141 OR L142) AND L144 AND (L147 AND
L142
L144
L147
L150
L172
                         L150)
=> s (1160 or 1163 or 1167 or 1168 or 1170-1172) not 1180
         13 (L160 OR L163 OR L167 OR L168 OR (L170 OR L171 OR L172)) NOT
L187
```

Searched by Barb O'Bryen, STIC 2-2518

L180 previously pinted

=> => dup rem 1183,1185,1182,1187,1184,1186 FILE 'MEDLINE' ENTERED AT 13:09:58 ON 03 FEB 2006

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FILE 'CAPLUS' ENTERED AT 13:09:58 ON 03 FEB 2006

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FILE 'WPIDS' ENTERED AT 13:09:58 ON 03 FEB 2006

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PROCESSING COMPLETED FOR L183 PROCESSING COMPLETED FOR L185 PROCESSING COMPLETED FOR L182 PROCESSING COMPLETED FOR L187 PROCESSING COMPLETED FOR L184 PROCESSING COMPLETED FOR L186

L188 63 DUP REM L183 L185 L182 L187 L184 L186 (6 DUPLICATES REMOVED)

> ANSWERS '1-18' FROM FILE MEDLINE ANSWERS '19-28' FROM FILE DRUGU ANSWERS '29-40' FROM FILE CAPLUS ANSWERS '41-52' FROM FILE BIOSIS ANSWERS '53-60' FROM FILE EMBASE ANSWERS '61-63' FROM FILE WPIDS

=> d iall 1-28; d ibib ed abs hitind 29-40; d iall 41-63; fil hom

L188 ANSWER 1 OF 63 MEDLINE on STN ACCESSION NUMBER: 1998371043 MEDLINE PubMed ID: 9705343

DOCUMENT NUMBER:

A novel carbohydrate-glycosphingolipid interaction between TITLE:

a beta-(1-3)-glucan immunomodulator, PGG-glucan, and

lactosylceramide of human leukocytes.

Zimmerman J W; Lindermuth J; Fish P A; Palace G P; AUTHOR:

Stevenson T T; DeMong D E

CORPORATE SOURCE: Alpha-Beta Technology, Inc., Worcester, Massachusetts

01605, USA.. jzimme@abti.com

SOURCE: Journal of biological chemistry, (1998 Aug 21) 273 (34)

22014-20.

Journal code: 2985121R. ISSN: 0021-9258.

PUB. COUNTRY:

United States Journal; Article; (JOURNAL ARTICLE)

DOCUMENT TYPE: LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199809

ENTRY DATE: Entered STN: 19980925

> Last Updated on STN: 19980925 Entered Medline: 19980917

ABSTRACT:

The immunomodulator Betafectin(R) PGG-glucan is a homopolymer of glucose

( roction

derived from yeast cell walls which has been demonstrated to enhance leukocyte anti-infective activity in vitro and in vivo, without the induction of proinflammatory cytokines. We report here the purification of a PGG-glucan-binding element from human leukocytes and its identification as lactosylceramide, a major glycosphingolipid of neutrophils, which includes the CDw17 epitope. The binding of radiolabeled PGG-glucan to purified lactosylceramide was saturable, specific, and time- and temperature-dependent. Lactosylceramides from human leukocytes were fractionated by high performance liquid chromatography in order to analyze the effect of ceramide structure on binding. A variety of fatty acid chain lengths with varying degrees of unsaturation were found to support binding to radiolabeled PGG-glucan. However, DL-lactosylceramides containing dihydrosphingosine did not bind. Radiolabeled PGG-glucan bound several other neutral glycosphingolipids with a terminal galactose, including galactosylceramide, globotriaosylceramide, and gangliotetraosylceramide. The binding of radiolabeled PGG-glucan to lactosylceramide was not inhibited by glycogen, dextran, mannan, pustulan, laminarin, or a low molecular weight beta-(1-3)-glucan, but was inhibited by high molecular weight beta-(1-3)-glucans and by a monoclonal antibody to lactosylceramide. Although this glycosphingolipid has been shown in numerous reports to bind various microorganisms, this represents the first report of lactosylceramide binding to a macromolecular carbohydrate.

CONTROLLED TERM: \*Adjuvants, Immunologic: ME, metabolism

Antigens, CD: ME, metabolism

Binding Sites

Cell Differentiation \*Glucans: ME, metabolism

\*Glycosphingolipids: ME, metabolism

\*Lactosylceramides: ME, metabolism

\*Leukocytes: ME, metabolism

Temperature Time Factors \*beta-Glucans

CAS REGISTRY NO.: 4682-48-8 (CDw17 antigen)

CHEMICAL NAME: 0 (Adjuvants, Immunologic); 0 (Antigens, CD); 0 (Glucans);

0 (Glycosphingolipids); 0 (Lactosylceramides); 0 (beta-Glucans); 0 (poly-1-6-glucopyranosyl-1-3-

glucopyranose glucan)

L188 ANSWER 2 OF 63 MEDLINE on STN ACCESSION NUMBER: 1999038741 MEDLINE PubMed ID: 9821300 DOCUMENT NUMBER:

Inhibition of lymphoproliferative response and its TITLE:

restoration with a glucan immunomodulator in mice with

experimental larval toxocarosis.

**AUTHOR:** Boroskova Z; Reiterova K; Dubinsky P; Tomasovicova O;

Machnicka B

Parasitological Institute, Slovak, Kosice, Slovakia. CORPORATE SOURCE:

Folia microbiologica, (1998) 43 (5) 475-6. Journal code: 0376757. ISSN: 0015-5632. SOURCE:

PUB. COUNTRY: Czech Republic

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199812

ENTRY DATE: Entered STN: 19990115

> Last Updated on STN: 19990115 Entered Medline: 19981211

ABSTRACT:

A high infective dose of Taxocara canis eggs (2,500 eggs per mouse) induced a

partial immunosuppression in mice, manifested by inhibition of the proliferative response of splenic T and B cells to polyclonal activators. A glucan immunomodulator given to infected animals at the beginning of the experiment showed a marked stimulative and restorative effect on the parasite-suppressed lymphoproliferative response. The ability of T. canis to migrate in the host was reduced in glucan-treated animals by 27%.

\*Adjuvants, Immunologic

Animals

CONTROLLED TERM:

B-Lymphocytes: IM, immunology Glucans: IM, immunology \*Glucans: PD, pharmacology

\*Immune Tolerance
Larva: IM, immunology
\*Lymphocyte Activation

Mice

Mice, Inbred C57BL

Mitogens: PD, pharmacology

Phytohemagglutinins: PD, pharmacology Research Support, Non-U.S. Gov't

T-Lymphocytes: IM, immunology
\*Toxocara canis: IM, immunology
Toxocara canis: PH, physiology
\*Toxocariasis: IM, immunology
Toxocariasis: PS, parasitology

\*beta-Glucans

CAS REGISTRY NO.: 9051-97-2 (beta-1,3-glucan)

CHEMICAL NAME: 0 (Adjuvants, Immunologic); 0 (Glucans); 0 (Mitogens); 0

(Phytohemagglutinins); 0 (beta-Glucans)

L188 ANSWER 3 OF 63 MEDLINE on STN
ACCESSION NUMBER: 1998351366 MEDLINE
DOCUMENT NUMBER: PubMed ID: 9688080

TITLE: Immunomodulatory effects of oat beta-glucan administered

intragastrically or parenterally on mice infected with

Eimeria vermiformis.

AUTHOR: Yun C H; Estrada A; Van Kessel A; Gajadhar A; Redmond M;

Laarveld B

CORPORATE SOURCE: Animal Biotechnology Centre, Department of Animal and

Poultry Science, University of Saskatchewan, Saskatoon,

Canada.

SOURCE: Microbiology and immunology, (1998) 42 (6) 457-65.

Journal code: 7703966. ISSN: 0385-5600.

PUB. COUNTRY: Japan

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199810

ENTRY DATE: Entered STN: 19981021

Last Updated on STN: 19981021 Entered Medline: 19981015

ABSTRACT:

The immunostimulatory effect of intragastrically or parenterally administered beta-(1-->3; 1-->4) glucan, extracted from oats (ObetaG), on disease resistance to Eimeria vermiformis was studied in C57BL/6 mice. Multiple administrations of ObetaG by intragastric or subcutaneous routes reduced fecal oocyst shedding compared to the non-treated control group. The administration of ObetaG by subcutaneous route resulted in higher levels of total serum immunoglobulins and antigen (sporozoite and merozoite)-specific immunoglobulins as compared with the non-treated group. To evaluate the effect of a single subcutaneous dose, groups of mice were treated with ObetaG 2 days before E. vermiformis infection,

at the time of infection and at 2 or 6 days after infection. From day 11 post-infection the oocyst discharge was significantly diminished (P<0.05-0.01) in the ObetaG-treated groups, except in those treated 6 days after infection, as compared to the non-treated control group. The proliferative responses to E. vermiformis sporozoite antigen of lymphocytes isolated from the spleen were significantly increased (P<0.05) when ObetaG was administered 2 days before or at the time of E. vermiformis infection. Lymphocyte proliferative responses to merozoite antigen were not influenced by treatment. In conclusion, ObetaG appeared to up-regulate immune mechanisms and provide enhanced resistance against eimerian coccidiosis in mice.

Check Tags: Female CONTROLLED TERM:

\*Adjuvants, Immunologic

Animals

Antibodies, Protozoan: BL, blood Antigens, Protozoan: IM, immunology

Avena sativa

\*Coccidiosis: IM, immunology Cytokines: BI, biosynthesis

\*Eimeria

Eimeria: GD, growth & development

Eimeria: IM, immunology

\*Glucans: AD, administration & dosage

\*Glucans: IM, immunology Immunoglobulins: BL, blood Lymphocyte Activation

Mice

Mice, Inbred C57BL

Research Support, Non-U.S. Gov't

Time Factors \*beta-Glucans

55965-23-6 (beta-glucan, (1-3)(1-4)-) CAS REGISTRY NO.:

0 (Adjuvants, Immunologic); 0 (Antibodies, Protozoan); 0 CHEMICAL NAME:

(Antigens, Protozoan); 0 (Cytokines); 0 (Glucans); 0

(Immunoglobulins); 0 (beta-Glucans)

L188 ANSWER 4 OF 63 MEDLINE on STN ACCESSION NUMBER: 1998151149 MEDLINE

PubMed ID: 9492185 DOCUMENT NUMBER:

Immunomodulatory activities of oat beta-glucan in vitro and TITLE:

Estrada A; Yun C H; Van Kessel A; Li B; Hauta S; Laarveld B AUTHOR:

Animal Biotechnology Centre, Department of Animal and CORPORATE SOURCE:

Poultry Science, University of Saskatchewan, Saskatoon,

Canada.. estrada@SASK.USASK.CA

Microbiology and immunology,  $(\underline{1997})$  41 (12) 991-8. Journal code: 7703966. ISSN:  $0\overline{385}$ -5600. SOURCE:

PUB. COUNTRY: Japan

Journal; Article; (JOURNAL ARTICLE) DOCUMENT TYPE:

LANGUAGE: English

Priority Journals FILE SEGMENT:

ENTRY MONTH: 199804

ENTRY DATE: Entered STN: 19980410

> Last Updated on STN: 19980410 Entered Medline: 19980402

## ABSTRACT:

Previous studies have shown that beta-qlucans extracted from yeast or fungi potentiate immune responses. In the present study, the immunomodulatory activities of beta-(1-->3,1-->4)-glucan, derived from oats, were investigated. The ability of oat beta-glucan (ObetaG) to stimulate\_IL-1 and TNF-alpha release from murine peritoneal macrophages and the murine macrophage cell line P338D1,

was assessed. In vitro stimulation of macrophages with ObetaG resulted in the production of IL-1 in a dose and time-dependent manner, whereas only small amounts of TNF-alpha could be detected in the culture supernatants. ObetaG also induced the production of IL-2, IFN-gamma and IL-4 secretion in a dose-dependent manner in cultured spleen cells. The intraperitoneal administration of ObetaG in mice resulted in the accumulation of leucocytes, predominantly macrophages, in the peritoneal cavity. Furthermore, ObetaG was tested for its ability to enhance non-specific resistance to a bacterial challenge in mice. Survival of mice challenged with Staphylococcus aureus was enhanced by a single intraperitoneal administration of 500 microg of ObetaG 3 days prior to bacterial challenge. In conclusion, these studies demonstrated that ObetaG possesses immunomodulatory activities capable of stimulating immune functions both in vitro and in vivo.

CONTROLLED TERM: \*Adjuvants, Immunologic: PD, pharmacology

Animals
Avena sativa
Cell Line
Cells, Cultured

\*Cytokines: ME, metabolism \*Glucans: IM, immunology

Interferon Type II: ME, metabolism

Interleukins: ME, metabolism

\*Macrophages, Peritoneal: IM, immunology

Mice

Mice, Inbred BALB C Mice, Inbred Strains

Peritoneal Cavity: CY, cytology Research Support, Non-U.S. Gov't

Spleen: IM, immunology

Staphylococcal Infections: IM, immunology Tumor Necrosis Factor-alpha: ME, metabolism

Zymosan: PD, pharmacology

\*beta-Glucans

CAS REGISTRY NO.: 55965-23-6 (beta-glucan, (1-3)(1-4)-); 82115-62-6

(Interferon Type II); 9010-72-4 (Zymosan)

CHEMICAL NAME: 0 (Adjuvants, Immunologic); 0 (Cytokines); 0 (Glucans); 0

(Interleukins); 0 (Tumor Necrosis Factor-alpha); 0

(beta-Glucans)

L188 ANSWER 5 OF 63 MEDLINE ON STN
ACCESSION NUMBER: 96229136 MEDLINE
DOCUMENT NUMBER: PubMod ID: 8644487

DOCUMENT NUMBER: PubMed ID: 8644497

TITLE: Glucans as immunological adjuvants.

AUTHOR: Mohagheghpour N; Dawson M; Hobbs P; Judd A; Winant R;

Dousman L; Waldeck N; Hokama L; Tuse D; Kos F; +

CORPORATE SOURCE: Life Sciences Division, SRI International, Menlo Park,

California 94025-3493, USA.

CONTRACT NUMBER: AI30939 (NIAID)

SOURCE: Advances in experimental medicine and biology,

13-22.

Journal code: 0121103. ISSN: 0065-2598.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199607

ENTRY DATE: Entered STN: 19960726

Last Updated on STN: 19960726 Entered Medline: 19960716

CONTROLLED TERM: Check Tags: Female

\*Adjuvants, Immunologic

Animals

Antibody Formation: DE, drug effects

Glucans: CH, chemistry \*Glucans: IM, immunology \*Glycoproteins: CH, chemistry Immunoconjugates: IM, immunology

Macrophages: IM, immunology

Mice

Mice, Inbred BALB C Molecular Structure

Rabbits

Research Support, U.S. Gov't, P.H.S.

\*Viral Proteins: IM, immunology

\*beta-Glucans

CAS REGISTRY NO.: 9051-97-2 (beta-1,3-glucan)

0 (Adjuvants, Immunologic); 0 (Glucans); 0 (Glycoproteins); CHEMICAL NAME:

0 (Immunoconjugates); 0 (Viral Proteins); 0 (beta-Glucans)

L188 ANSWER 6 OF 63 MEDLINE on STN ACCESSION NUMBER: 94176560 MEDLINE DOCUMENT NUMBER: PubMed ID: 8130277

TITLE: A novel immunomodulator soluble aminated beta-1.3-D-glucan:

binding characteristics to mouse peritoneal macrophages. Konopski Z; Smedsrod B; Seljelid R; Eskeland T

AUTHOR: CORPORATE SOURCE:

Department of Experimental Pathology and Anatomy,

University of Tromso, Norway.

Biochimica et biophysica acta, (1994 Mar 10) 1221 (1) 61-5. SOURCE:

Journal code: 0217513. ISSN: 0006-3002.

PUB. COUNTRY: Netherlands

Journal; Article; (JOURNAL ARTICLE) DOCUMENT TYPE:

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199404

ENTRY DATE: Entered STN: 19940428

> Last Updated on STN: 19970203 Entered Medline: 19940418

#### ABSTRACT:

We have previously reported that soluble aminated beta-1.3-D-glucan (AG), a potent immunomodulator, specifically inhibited binding and internalization of AG-coated microbeads (GDM) in mouse peritoneal macrophages. The present study was undertaken to determine parameters of AG binding to macrophages. For this purpose, AG was conjugated with tyraminyl cellobiose (TC), which can be radioiodinated. With this method the immunomodulator was labelled with a very high specific radioactivity, allowing sensitive measurements of binding. Maximal binding capacity was 0.33 micrograms [1251]TC-AG/10(6) cells. Binding was inhibited by TC-AG and AG, but not by mannose and mannan, showing that the receptor different from the mannose receptor was involved. Binding was reversible, with an initial association rate of 120 cpm/min, and a much faster initial dissociation rate of 680 cpm/min. Bound [1251]TC-AG was internalized. These findings suggest that both AG and GDM are bound and internalized via the same beta-glucan receptor in mouse peritoneal macrophages.

Check Tags: Female CONTROLLED TERM:

\*Adjuvants, Immunologic: ME, metabolism

Animals

Biological Transport Cells, Cultured

\*Glucans: ME, metabolism

Iodine Radioisotopes

Kinetics

16 (4)

\*Macrophages, Peritoneal: ME, metabolism

Mice

Mice, Inbred BALB C Mice, Inbred C57BL

\*beta-Glucans

CAS REGISTRY NO.:

9051-97-2 (beta-1,3-glucan)

CHEMICAL NAME:

0 (Adjuvants, Immunologic); 0 (Glucans); 0 (Iodine

Radioisotopes); 0 (beta-Glucans)

L188 ANSWER 7 OF 63 MEDLINE ON STN ACCESSION NUMBER: 93364375 MEDLINE DOCUMENT NUMBER: PubMed ID: 8358393

TITLE:

Immunopharmacological characterization of a highly branched fungal (1-->3)-beta-D-glucan, OL-2, isolated from Omphalia

lapidescens.

AUTHOR:

Ohno N; Saito K; Nemoto J; Kaneko S; Adachi Y; Nishijima M;

Miyazaki T; Yadomae T

CORPORATE SOURCE:

Tokyo College of Pharmacy, Japan.

SOURCE:

Biological & pharmaceutical bulletin, (1993 Apr)

414-9.

Journal code: 9311984. ISSN: 0918-6158.

PUB. COUNTRY:

Japan

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

199309

ENTRY DATE:

Entered STN: 19931015

Last Updated on STN: 19931015 Entered Medline: 19930927

## ABSTRACT:

The immunopharmacological activities of a fungal (1-->3)-beta-D-glucan, OL-2, isolated from "Leiwan" Omphalia lapidescens were examined. Intraperitoneal (i.p.) administration of OL-2 to ICR mice induced a significant number of peritoneal exudate cells (PEC) and white blood cells over the period of a few days. Spleen cell numbers were also increased by i.p. administration of OL-2 at about a week. These changes reverted to the normal level within a month. Responses of spleen cells and bone marrow cells (BM) to colony stimulating factors (CSF) were augmented by OL-2 administration assessed by cell proliferation assay. Sera from OL-2 administered mice contained an increased concentration of colony stimulating activity. Gene expressions of interleukin-1 beta, interleukin-6, and tumor necrosis factor alpha in the spleen were also increased. These results suggested the activation of hematopoietic responses, and would well relate to the incremental increase in PEC, white blood cell and spleen cell numbers. OL-2 also increased the serum concentration of fibronectin and complement component C-3. However, OL-2 did not show adjuvant activity to SRBC and antitumor activity against the solid form of Sarcoma 180 by i.p. administration. Yet, OL-2 did not interfere with the antitumor activity of SSG against the same tumor system. These facts suggested that OL-2 could enhance nonspecific host defense mechanisms by enhancing hematopoietic responses, but would not enhance or inhibit the specific immunity mediated by lymphocytes. (ABSTRACT TRUNCATED AT 250 WORDS) Check Tags: Male CONTROLLED TERM:

Adjuvants, Immunologic: PD, pharmacology

\*Agaricales: CH, chemistry

Animals

Antineoplastic Agents: CH, chemistry Antineoplastic Agents: IM, immunology \*Antineoplastic Agents: PD, pharmacology

Base Sequence

Chemistry, Physical

Complement 3: BI, biosynthesis Cytokines: BI, biosynthesis Fibronectins: BL, blood Fibronectins: IM, immunology Glucans: CH, chemistry

Glucans: IM, immunology \*Glucans: PD, pharmacology

Leukocyte Count: DE, drug effects Leukocytes: DE, drug effects Leukocytes: IM, immunology

Lymphocyte Activation: DE, drug effects

Mice

Mice, Inbred AKR Mice, Inbred ICR

Molecular Sequence Data

RNA, Messenger: BI, biosynthesis Sarcoma 180: DT, drug therapy Structure-Activity Relationship

CAS REGISTRY NO.: 96778-06-2 (OL 2)

0 (Adjuvants, Immunologic); 0 (Antineoplastic Agents); 0 CHEMICAL NAME:

(Complement 3); 0 (Cytokines); 0 (Fibronectins); 0

(Glucans); 0 (RNA, Messenger)

MEDLINE on STN L188 ANSWER 8 OF 63 ACCESSION NUMBER: 92380573 MEDLINE PubMed ID: 1823656 DOCUMENT NUMBER:

The immunoadjuvant effect of soluble glucan derivatives in TITLE:

Wagnerova J; Liskova A; Cervenakova L; Trnovec T; Ferencik AUTHOR:

CORPORATE SOURCE: Institute of Experimental Pharmacology, Slovak Academy of

Sciences, Bratislava, Czechoslovakia.

Folia microbiologica, (1991) 36 (2) 198-204. Journal code: 0376757. ISSN: 0015-5632. SOURCE:

PUB. COUNTRY: Czechoslovakia

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199209

ENTRY DATE: Entered STN: 19921018

> Last Updated on STN: 19921018 Entered Medline: 19920928

## ABSTRACT:

We examined the effect of soluble derivatives of yeast glucan on the humoral immune response of various strains of inbred mice after administration of different doses according to various schedules. Glucan was injected i.v. or s.c. in a single dose or repeatedly. The immune response was examined by determining the titres of serum hemagglutinins against sheep erythrocytes (SRBC-Ab). The immunoadjuvant effect of glucan derivatives depends on the inbred strain used, on the dose of glucan, mode and time of administration with respect to antigen injection. The results have shown that the stimulatory effect of glucan derivatives occurred already after a single injection, the optimum dose being 10-20 mg/kg. Intravenous injection was more efficient than the subcutaneous one. In some cases, a slight increase of the spleen mass was observed.

CONTROLLED TERM: Check Tags: Female

Adjuvants, Immunologic: AD, administration &

\*Adjuvants, Immunologic: PD, pharmacology

Animals

combination in

Antibody Formation

Dose-Response Relationship, Immunologic Glucans: AD, administration & dosage

\*Glucans: IM, immunology Glucans: PD, pharmacology

\*Hemagglutinins: BI, biosynthesis

Mice

Mice, Inbred A Mice, Inbred C57BL Mice, Inbred CBA

Sheep

Species Specificity Yeasts: CH, chemistry Yeasts: IM, immunology

0 (Adjuvants, Immunologic); 0 (Glucans); 0 (Hemagglutinins) CHEMICAL NAME:

MEDLINE on STN L188 ANSWER 9 OF 63 90152775 MEDLINE ACCESSION NUMBER:

DOCUMENT NUMBER:

PubMed ID: 2695459

Immunoprotection by beta-1,3 glucan antigen TITLE:

Plasmodium berghei infection in mice.

**AUTHOR:** Maheshwari R; Siddiqui M U

Indian journal of medical research, SOURCE:

((1989 Nov)

Journal code: 0374701. ISSN: 0971-591⁄6

PUB. COUNTRY: India

Journal; Article; (JOURNAL ARTICLE) DOCUMENT TYPE:

English LANGUAGE:

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199003

ENTRY DATE: Entered STN: 19900601

> Last Updated on STN: 19900601 Entered Medline: 19900321

#### ABSTRACT:

In an attempt to protect mice against experimental infection with P. berghei, mice were immunized against soluble extract of P. berghei in combination with beta-1,3 glucan or FCA and also independently. Mice immunized against P. berghei antigen-glucan developed well defined cell mediated and humoral immune responses, while mice injected with antigen FCA or antigen alone developed only an antibody response. Antigen-glucan immunization afforded a high degree of immune protection to the host against the challenge with live parasites.

Check Tags: Male CONTROLLED TERM:

\*Adjuvants, Immunologic

Animals

Antibodies, Protozoan: BI, biosynthesis \*Antigens, Protozoan: IM, immunology

\*Glucans: IM, immunology

Immunity, Cellular

\*Malaria: PC, prevention & control

Mice

\*Plasmodium berghei: IM, immunology

\*beta-Glucans

9051-97-2 (beta-1,3-glucan) CAS REGISTRY NO.:

CHEMICAL NAME: 0 (Adjuvants, Immunologic); 0 (Antibodies, Protozoan); 0 (Antigens, Protozoan); 0 (Glucans); 0 (beta-Glucans)

MEDLINE on STN L188 ANSWER 10 OF 63 MEDLINE ACCESSION NUMBER: 89291140 DOCUMENT NUMBER: PubMed ID: 2737802

TITLE: Protective effect of L. donovani antigens using glucan as

an adjuvant.

Jones 10/630143

AUTHOR: Obaid K A; Ahmad S; Khan H M; Mahdi A A; Khanna R CORPORATE SOURCE: Department of Microbiology, Jawaharlal Nehru Medical

College, Aligarh Muslim University, India.

SOURCE: International journal of immunopharmacology, (1989) 11 (3)

229-35.

Journal code: 7904799. ISSN: 0192-0561.

PUB. COUNTRY: ENGLAND: United Kingdom

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 198908

ENTRY DATE: Entered STN: 19900309

Last Updated on STN: 19900309 Entered Medline: 19890803

## ABSTRACT:

Golden hamsters were immunized with various antigen fractions of Leishmania donovani promastigotes. Beta 1,3-glucan was used as an adjuvant in these vaccination experiments. The results indicate that immunization of animals with the microsomal fraction (subcellular fraction III) in combination with glucan confers considerable immune protection against L. donovani infection. The immune protection was confirmed by correspondingly lower parasite burden in the livers and spleens of test animals compared to controls. Additionally, the vaccinated animals showed positive skin test responsiveness after challenge, along with increased antibody titres. Immunization of animals with whole and particulate antigen fractions was also found to afford a high degree of resistance. The other subcellular and soluble antigen fractions conferred very little protection. In these experiments, glucan was found to be a potent adjuvant when injected, intraperitoneally, with Leishmania antigens. Similar doses of parasite extracts given without an adjuvant were able to confer only very little or no protection.

CONTROLLED TERM: Check Tags: Male

\*Adjuvants, Immunologic

Animals

Antibody Formation

\*Antigens, Protozoan: IM, immunology Enzyme-Linked Immunosorbent Assay

\*Glucans: IM, immunology

Hamsters

Immunity, Cellular

\*Leishmania donovani: IM, immunology

Liver: PS, parasitology Liver: PA, pathology Spleen: PA, pathology

CHEMICAL NAME: 0 (Adjuvants, Immunologic); 0 (Antigens, Protozoan); 0

(Glucans)

L188 ANSWER 11 OF 63 MEDLINE ON STN ACCESSION NUMBER: 86053262 MEDLINE DOCUMENT NUMBER: PubMed ID: 3079591

TITLE: Calcium-dependent and -independent tumoricidal activities

of polymorphonuclear leukocytes induced by a linear

beta-1,3-D-glucan and phorbol myristate acetate in mice.
Morikawa K; Noguchi T; Yamazaki M; Mizuno D

AUTHOR: Morikawa K; Noguchi T; Yamazaki M; Mizuno E SOURCE: Cancer research, (1986 Jan) 46 (1) 66-70.

Journal code: 2984705R. ISSN: 0008-5472.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 198601

ENTRY DATE:

CONTROLLED TERM:

CAS REGISTRY NO.:

Entered STN: 19900321

Last Updated on STN: 19970203 Entered Medline: 19860123

ABSTRACT:

Some antitumor immunomodulators, such as a linear beta-1,3-D-glucan from Alcaligenes faecalis var. myxogenes IFO 13140 (TAK), induce potent tumoricidal activity of polymorphonuclear leukocytes (PMNs). In the present study we investigated the role of calcium on the tumoricidal activity of PMNs induced by immunomodulators, especially TAK. The calcium chelator ethylene glycol bis (beta-aminoethyl ether) -N, N, N', N'-tetraacetic acid (EGTA) almost completely inhibited TAK-induced PMN cytotoxicity and this inhibition was restored by Ca2+ but not by Mg2+. In Ca2+- and Mg2+-free medium, PMN cytotoxicity induced by TAK was recovered by the addition of Ca2+ provided that Mg2+ was also present. By scopoletin assay, hydrogen peroxide released from PMNs by TAK was also observed in the presence of Ca2+ but not in its absence. The PMN cytotoxicities induced by the other immunomodulators, Propionibacterium acnes, Bacillus Calmette-Guerin, zymosan A, and Nocardia cell wall skeletons were also Ca2+ dependent, judging from studies with EGTA and measurement of hydrogen peroxide release in the presence and absence of Ca2+. The Ca2+ dependency of these PMN cytotoxicities suggests that Ca2+ influx is involved in the cytolytic process, but PMN cytotoxicity was not induced by simple addition of the calcium ionophore A23187. Like TAK, phorbol myristate acetate induced PMN cytotoxicity but this cytotoxicity was not Ca2+ dependent. The present report demonstrates the difference in Ca2+ dependency of these PMN cytotoxicities; i.e., extracellular calcium was required for immunomodulator-induced PMN cytotoxicity, but not for phorbol myristate acetate-induced PMN cytotoxicity. This suggests that the processes of induction of PMN cytotoxicity by the two types of activators are not identical.

\*Adjuvants, Immunologic: IM, immunology

Animals

Check Tags: Male

Calcimycin: PD, pharmacology

\*Calcium: PH, physiology

Cytotoxicity, Immunologic: DE, drug effects

Egtazic Acid: PD, pharmacology

\*Glucans: IM, immunology

Hydrogen Peroxide: ME, metabolism

Magnesium: PD, pharmacology

Mice

Mice, Inbred C3H

\*Neutrophils: IM, immunology \*Phorbols: PD, pharmacology

Research Support, Non-U.S. Gov't

\*Tetradecanoylphorbol Acetate: PD, pharmacology 16561-29-8 (Tetradecanoylphorbol Acetate); 52665-69-7

(Calcimycin); 67-42-5 (Egtazic Acid); 7439-95-4

(Magnesium); 7440-70-2 (Calcium); 7722-84-1 (Hydrogen

Peroxide)

CHEMICAL NAME: 0 (Adjuvants, Immunologic); 0 (Glucans); 0 (Phorbols)

L188 ANSWER 12 OF 63 MEDLINE ON STN ACCESSION NUMBER: 85291288 MEDLINE DOCUMENT NUMBER: PubMed ID: 3861748

TITLE: Preparation for hapten help by glucan, muramyl dipeptide,

and its L-ala-Glycerol-mycolate derivative.

AUTHOR: Leech S H; Di Luzio N R; Leclerc C

CONTRACT NUMBER: CA 24326 (NCI)

CA 25668 (NCI)

SOURCE: Journal of leukocyte biology, (1985 Aug) 38 (2) 317-25.

Journal code: 8405628. ISSN: 0741-5400.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 198510

ENTRY DATE: Entered STN: 19900320

Last Updated on STN: 19970203 Entered Medline: 19851004

#### ABSTRACT:

Previously, we reported that one of the factors that determines whether or n∮t an animal will be prepared for hapten help after priming is the type of adjuvant used. The present work was undertaken, therefore, to determine whach of a diverse variety of adjuvants or biological response modifiers would be effective. They included Freund's complete (CFA) and incomplete (FICA) adjuvants, particulate glucan, muramyl dipeptide (MDP), and its L-ala-glycerol-mycolate derivative. Help by the azobenzenearsonate (ABA) hapten was measured as the augmentation of the anti-bovine gamma-globulin (BGG) plaque-forming cell (PFC) response to ABA-BGG of mice that had been hapten-primed with ABA conjugated to ovalbumin (OVA). The results showed that FICA was ineffective. MDP was effective but only if administered with FICA during hapten-priming. MDP-L-ala-glycerol-mycolate was effective without any adjuvant but only within a narrow dose range. Particulate glucan was as effective as CFA in preparing mice for hapten help. As the macrophage is the primary cellular target of those biological response modifiers that were effective, we conclude that it plays an important role in the cellular interaction involved in the mediation of hapten help.

CONTROLLED TERM: \*Acetylmuramyl-Alanyl-Isoglutamine: AA, analogs &

derivatives

\*Acetylmuramyl-Alanyl-Isoglutamine: IM, immunology

\*Adjuvants, Immunologic

Animals

Antibody Formation

\*Glucans: IM, immunology

\*Haptens: IM, immunology Lymphocytes: IM, immunology \*Macrophages: IM, immunology

Mice

Mice, Inbred BALB C

Research Support, Non-U.S. Gov't Research Support, U.S. Gov't, P.H.S.

CAS REGISTRY NO.: 53678-77-6 (Acetylmuramyl-Alanyl-Isoglutamine)

CHEMICAL NAME: 0 (1-0-(acetylmuramyl-alanyl-isoglutaminyl-alanine)-

glycerol-3-mycolate); 0 (Adjuvants, Immunologic); 0

(Glucans); 0 (Haptens)

L188 ANSWER 13 OF 63 MEDLINE ON STN ACCESSION NUMBER: 86157880 MEDLINE DOCUMENT NUMBER: PubMed ID: 3913389

TITLE: Glucan-induced immunity in mice against Plasmodium berghei.

AUTHOR: Kumar P; Ahmad S

SOURCE: Annals of tropical medicine and parasitology, (1985 Apr) 79

(2) 211-3.

Journal code: 2985178R. ISSN: 0003-4983.

PUB. COUNTRY: ENGLAND: United Kingdom

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 198604

ENTRY DATE: Entered STN: 19900321

Last Updated on STN: 19900321

6 (5)

Entered Medline: 19860411

CONTROLLED TERM:

\*Adjuvants, Immunologic

Animals

\*Antibody Formation

Antigens, Protozoan: IM, immunology

Cell Migration Inhibition

Glucans: BL, blood

\*Glucans: IM, immunology

Immunization

Mice

\*Plasmodium berghei: IM, immunology Research Support, Non-U.S. Gov't

\*beta-Glucans

CAS REGISTRY NO.: 9051-97-2 (beta-1,3-glucan)

CHEMICAL NAME: 0 (Adjuvants, Immunologic); 0 (Antigens, Protozoan); 0

(Glucans); 0 (beta-Glucans)

L188 ANSWER 14 OF 63 MEDLINE ON STN ACCESSION NUMBER: 85053635 MEDLINE DOCUMENT NUMBER: PubMed ID: 6094370

TITLE: Immunization of guinea pigs against Entamoeba histolytica

using glucan as an adjuvant.

AUTHOR: Sharma A; Haq A U; Siddiqui M U; Ahmad S

SOURCE: International journal of immunopharmacology,

483-91.

Journal code: 7904799. ISSN: 0192-0561.

PUB. COUNTRY: ENGLAND: United Kingdom

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 198412

ENTRY DATE: Entered STN: 19900320

Last Updated on STN: 19900320 Entered Medline: 19841226

ABSTRACT:

Beta 1-3 polyglucose or glucan, an extract of cell wall of Saccharomyces cerevisiae, has been successfully employed in this laboratory as an effective immunopotentiator in experimental studies on amoebiasis. An antigen extract from Entamoeba histolytica was combined with beta, 1-3 glucan for immunizing guinea pigs. In order to study the effectiveness of such vaccine preparations, several batches of guinea pigs were immunized with amoeba antigen alone, and in combination with various immunoadjuvants. Antigen inoculations were carried out via intraperitoneal route. Protective immune responses were obtained against amoeba antigen by using glucan as an adjuvant partner. The study showed that glucan can be safely used as an effective immune enhancer.

CONTROLLED TERM: \*Adjuvants, Immunologic

Animals

\*Entamoeba histolytica: IM, immunology

\*Glucans: IM, immunology

Guinea Pigs

Hemagglutination Tests: MT, methods Research Support, Non-U.S. Gov't

Skin Tests

\*Vaccines: IM, immunology

\*beta-Glucans

CAS REGISTRY NO.: 9051-97-2 (beta-1,3-glucan)

CHEMICAL NAME: 0 (Adjuvants, Immunologic); 0 (Glucans); 0 (Vaccines); 0

(beta-Glucans)

L188 ANSWER 15 OF 63 MEDLINE on STN

Jones 10/630143

Page 40

ACCESSION NUMBER: 83212008 MEDLINE DOCUMENT NUMBER: PubMed ID: 6852909

TITLE: Immunogenicity of soluble and particulate antigens from

Leishmania donovani: effect of glucan as an adjuvant.

AUTHOR: Cook J A; Holbrook T W

CONTRACT NUMBER: AI-18039 (NIAID)

SOURCE: Infection and immunity, (1983 Jun) 40 (3) 1038-43.

Journal code: 0246127. ISSN: 0019-9567.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTÍCLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 198307

ENTRY DATE: Entered STN: 19900319

Last Updated on STN: 19970203 Entered Medline: 19830715

## ABSTRACT:

The protective efficacy of glucan as an adjuvant with killed promastigotes of Leishmania donovani was compared with that of soluble or particulate fractions of the parasite. When these vaccine preparations were injected either intravenously or subcutaneously in CF-1 mice, glucan potentiated resistant against L. donovani infections as reflected by significant reductions in hepatic amastigote counts relative to infected control mice. The leishmanial antigens alone afforded no protection. Serum direct agglutination titexs to leishmanial antigens were highest in all groups given the vaccine intravenously, whereas the delayed-type hypersensitivity response to the antigen was positive only in groups immunized subcutaneously with glu $\!\!\!/ \epsilon$ an as an adjuvant. Some index of protection and immune response against visc@ral infection with the parasite was seen in groups vaccinated with gluca and soluble antigens. However, the protection afforded by glucan and particulate antigens of L. donovani more closely paralleled the resistance of wice treated with glucan and unfractionated killed promastigotes. Further antigenic analysis of particulate fractions of L. donovani may optimize effetive immunization when used with appropriate adjuvants, e.g., glucan.

CONTROLLED TERM: Check Tags: Comparative Study; Female

\*Adjuvants, Immunologic
Agglutinins: AN, analysis

Animals

\*Glucans: IM, immunology
Hypersensitivity, Delayed

\*Immunization

Injections, Intravenous Injections, Subcutaneous

Leishmania: GD, growth & development

\*Leishmania: IM, immunology

\*Leishmaniasis, Visceral: IM, immunology Leishmaniasis, Visceral: PS, parasitology

Liver: PS, parasitology

Mice

Research Support, U.S. Gov't, P.H.S.

Solubility

CHEMICAL NAME: 0 (Adjuvants, Immunologic); 0 (Agglutinins); 0 (Glucans)

L188 ANSWER 16 OF 63 MEDLINE ON STN ACCESSION NUMBER: 82166374 MEDLINE DOCUMENT NUMBER: PubMed ID: 7068224

TITLE: Glucan as an adjuvant for a murine Babesia microti

immunization trial.

AUTHOR: Benach J L; Habicht G S; Holbrook T W; Cook J A

CONTRACT NUMBER: AG00801 (NIA)

Jones 10/630143 Page 41

SOURCE:

Infection and immunity, ((1982 Mar)) 35 (3) 947-51.

Journal code: 0246127. ISSN: 001/9-9567.

PUB. COUNTRY:

United States

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

198206

ENTRY DATE:

Entered STN: 19900317

Last Updated on STN: 19970203 Entered Medline: 19820614

#### ABSTRACT:

A vaccination protocol against murine Babesia microti infection, using glucan, a beta-1.3-glucopyranose derivative of yeast cell walls, and qlutaraldehyde-fixed infected erythrocytes was evaluated. BALB/c mice were immunized intravenously four times at 2-day intervals with 2 X 10(8) fixed infected erythrocytes with and without glucan (0.45 mg). The mice were challenged 2 weeks after the last immunization with 10(8) viable infected erythrocytes. Peak parasitemia was significantly reduced (8.9 +/- 1.0%; P less than 0.001) in glucan-immunized mice as compared with nonimmunized controls (41.2 +/- 1.4%), glucan-treated controls (31.7 +/- 2.5%); P less than (0.05), or mice which received fixed infected erythrocytes without glucan (21.0 +/- 1.2%; P less than 0.01). Humoral and cellular immunity to B. microti was evaluated  $\overline{\phantom{a}}$ before challenge by measuring antibody titers and splenocyte blastogenic responses to B. microti antigens. The in vitro cellular response was inversely correlated with mean peak parasitemia (P less than 0.05). These observations demonstrate that glucan is an effective adjuvant in enhancing immunity to murine babesiosis.

CONTROLLED TERM:

\*Adjuvants, Immunologic

Animals

Antibodies: AN, analysis
\*Babesia: IM, immunology
\*Babesiosis: IM, immunology
Drug Evaluation, Preclinical
\*Glucans: IM, immunology
Lymphocyte Activation

Mice

Mice, Inbred BALB C.

Research Support, Non-U.S. Gov't Research Support, U.S. Gov't, P.H.S.

Vaccination

\*Vaccines: IM, immunology

CHEMICAL NAME:

0 (Adjuvants, Immunologic); 0 (Antibodies); 0 (Glucans); 0

(Vaccines)

L188 ANSWER 17 OF 63

MEDLINE on STN 82277650 MEDLINE

ACCESSION NUMBER: DOCUMENT NUMBER:

PubMed ID: 7051368

TITLE:

Immunopotentiation of anticancer chemotherapy by Candida

albicans, other yeasts and insoluble glucan in an

experimental lymphoma model.

AUTHOR:

Cassone A; Bistoñi F; Cenci E; Pesce C D; Tissi L; Marconi

Р

SOURCE:

Sabouraudia, (1982 Jun.) 20 (2) 115-25. Journal code: 0417341. ISSN: 0036-2174.

PUB. COUNTRY:

SCOTLAND: United Kingdom

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

198210

ENTRY DATE:

Entered STN: 19900317

Last Updated on STN: 19900317 Entered Medline: 19821021

#### ABSTRACT:

Several yeast species in the genera Candida, Saccharomyces and Cryptococcus showed powerful immunoadjuvant, chemotherapy-synergic effects against a histocompatible, virus-induced murine lymphoma. Sensitizing and booster intraperitoneal injections of 2 x 10(7) yeast cells on days -14 and +1 (with respect to tumor challenge on day 0) followed by treatment with antiblastic drugs (on day +5) were required to elicit optimum activity. The antitumor effect was not markedly influenced by the morphological growth form of merthiolate-inactivated C. albicans nor by the nature of the carbon source in the growth medium, except for C. albicans cells grown in a medium containing stearic acid, which were not effective. These cells had a higher ratio of soluble to insoluble cell wall components, as compared to glucose-grown cells, but this finding alone could hardly explain the lack of antitumor effects. Previous observations, suggesting that the alkali-acid insoluble beta-glucan (in the form of cell wall ghosts) is the only component of yeast cell walls endowed with antitumor activity comparable to that of whole cells, were confirmed and extended; the soluble mannan and glucan-protein fractions were unable to replace whole cells and glucan ghosts even as sensitizers or as boosting agents.

CONTROLLED TERM: \*Adjuvants, Immunologic: TU, therapeutic use

Animals

\*Antineoplastic Agents: TU, therapeutic use

Candida albicans: AN, analysis Candida albicans: CY, cytology Candida albicans: IM, immunology Carmustine: TU, therapeutic use

Cell Wall: AN, analysis Cryptococcus: IM, immunology

Culture Media

Fluorouracil: TU, therapeutic use

\*Glucans: IM, immunology \*Lymphoma: TH, therapy Mice

Neoplasms, Experimental: TH, therapy Research Support, Non-U.S. Gov't

Saccharomyces

\*Yeasts: IM, immunology

154-93-8 (Carmustine); 51-21-8 (Fluorouracil) CAS REGISTRY NO.:

0 (Adjuvants, Immunologic); 0 (Antineoplastic Agents); 0 CHEMICAL NAME:

(Culture Media); 0 (Glucans)

MEDLINE on STN L188 ANSWER 18 OF 63 ACCESSION NUMBER: 81238596 MEDLINE PubMed ID: 7019074 DOCUMENT NUMBER:

Glucan-enhanced immunogenicity of killed erythrocyte stages TITLE:

of Plasmodium berghei.

Holbrook T W; Cook J A; Parker B W AUTHOR:

Infection and immunity, (1981 May) 32 (2) 542-6. Journal code: 0246127. ISSN: 0019-9567. SOURCE:

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

Priority Journals FILE SEGMENT:

ENTRY MONTH: 198109

ENTRY DATE: Entered STN: 19900316

> Last Updated on STN: 19970203 Entered Medline: 19810915

ABSTRACT:

Intravenous injections of glucan simultaneously with Formalin-killed erythrocytic stages of Plasmodium berghei elicited a greater degree of resistance in mice against subsequent infection with viable parasites than injections of killed erythrocytic stages alone. In two experiments with P. berghei strain NK 65, 100% of mice immunized with the glucan-dead parasite preparation survived challenge, whereas only 28.6% of mice receiving dead parasites alone survived. In the third experiment, using P. berghei strain NYU-2, the same proportion of mice survived after immunization with glucan and dead parasites as with dead parasites alone (i.e., 10 of 11 in each group), but mice immunized with the glucan-dead parasite preparation experienced parasitemias of significantly less intensity and shorter duration than mice which received only dead parasites before infection. Inoculation of glucan alone or with normal erythrocytes conferred no protection against challenge.

CONTROLLED TERM: \*Adjuvants, Immunologic

Animals

Erythrocytes: PS, parasitology

\*Glucans: IM, immunology

Immunization

Immunization Schedule
\*Malaria: IM, immunology
Malaria: PS, parasitology

Mice

\*Plasmodium berghei: IM, immunology

CHEMICAL NAME: 0 (Adjuvants, Immunologic); 0 (Glucans)

L188 ANSWER 19 OF 63 DRUGU COPYRIGHT 2006 THE THOMSON CORP on STN DUPLICATE

ACCESSION NUMBER: 1989-45413 DRUGU P

TITLE: Immunomodulation by Orally Administered beta-Glucan in Mice.

AUTHOR: Suzuki I; Hashimoto K; Ohno N; Tanaka H; Yadomae T

LOCATION: Tokyo, Japan

SOURCE: Int.J.Immunopharmacol. (11, No. 7, 761-69, 1989) 3 Fig. 3

Tab. 36 Ref.

CODEN: IJIMDS ISSN: 0192-0561

AVAIL. OF DOC.: Laboratory of Immunopharmacology of Microbial Products, Tokyo

College of Pharmacy, Horinouchi 1432-1, Hachioji, Tokyo

192-03, Japan.

LANGUAGE: English DOCUMENT TYPE: Journal

ABSTRACT:

SSG, a beta-1,3-D-glucan, given p.o. to mice, augmented the proliferative response of spleen cells to Con-A or lipopolysaccharide (LPS).P.o.SSG also enhanced the activities of both natural killer (NK) cells in spleen and lysosomal enzymes of peritoneal macrophages. Augmentation of NK cells by SSG was less than that of poly I:C (Yamasa-Shoyu). SSG also possessed antitumor effects in IMC carcinoma and Meth A fibrosarcoma-bearing mice. A more purified SASG (P-SSG) produced similar effects. Results show that p.o. SSG can potentiate the immune response of mice.

SECTION HEADING: P Pharmacology

CLASSIF. CODE: 20 Immunological

50 Biological Response Modifiers
52 Chemotherapy - non-clinical

CONTROLLED TERM:

[01] GLUCAN-BETA-1,3-D \*PH; METH-A \*OC; FIBROSARCOMA

\*OC; ANIMAL-NEOPLASM \*OC; CARCINOMA \*OC; CONCANAVALIN-A \*RC;

ENDOTOXIN \*RC; POLY-I-C \*RC; YAMASA-SHOYU \*FT;

NAT.KILLER-CELL \*FT; P.O. \*FT; SPLEEN-CELL \*FT; MACROPHAGE

\*FT; CYTOSTATIC \*FT; MOUSE \*FT; IN-VIVO \*FT; IMMUNOSTIMULANT \*FT; IMMUNE-RESPONSE \*FT; TOXINS \*FT; LYMPHOCYTE \*FT; RES \*FT; LAB.ANIMAL

\*FT; IMMUNITY \*FT; GLUCAB13D \*RN; PH \*FT

FIELD AVAIL.: AB; LA; CT FILE SEGMENT: Literature

L188 ANSWER 20 OF 63 DRUGU COPYRIGHT 2006 THE THOMSON CORP on STN

ACCESSION NUMBER: 2000-01820 DRUGU P

TITLE: Comparison of the effect of different immunological adjuvants

on the antibody and T cell response to immunization with

MUC1-KLH and GD3-KLH conjugate cancer vaccines.

AUTHOR: Kim S K; Ragupathi G; Musselli C; Choi S J; Park Y S;

Livingston P O

CORPORATE SOURCE: Memorial-Sloan-Kettering-Cancer-Cent.; Univ.Yonsei;

Univ.Kwandong

LOCATION: New York, N.Y., USA; Wonju; Kangnung, South Korea SOURCE: Vaccine (18, No. 7-8, 597-603, 1999) 6 Tab. 20 Ref.

CODEN: VACCDE ISSN: 0264-410X

AVAIL. OF DOC.: Laboratory of Developmental Tumor Vaccinology, Memorial

Sloan-Kettering Cancer Center, 1275 York Avenue, New York, NY

10021, U.S.A. (P.O.L.).

LANGUAGE: English DOCUMENT TYPE: Journal

## ABSTRACT:

19 Immunological adjuvants (Adjuvax, Betafectin GG-glucan (both Alpha-Beta-Technologies), CRL-1005 (Vaxcel), CpG ODN (CpG pharmaceutics), DTP-N-GDP (amide/dextrose) (Endorex), Detox-PC, MPL-SE (both Ribi), GERBU adjuvant 10 (Biotech-Corp), MoGM-CSF (Lexigen), PSC97B adjuvant (Protein Sci.), QS-21 (Aquila), TiterMax Gold (CytRx), Adjumer, beta-alethine +/-peptide, GSK-1, GcMAF, MPC-026, PG-026) were compared for potentiation of spleen lymphocyte proliferation, cytokine release and antibody responses after s.c. immunization with keyhole limpet hemocyanin (KLH) conjugates with the human cancer antigens MUC1 peptide and GD3 ganglioside. Overall, QS-1 was the most effective.

SECTION HEADING: P Pharmacology

CLASSIF. CODE: 20 Immunological

50 Biological Response Modifiers 52 Chemotherapy - non-clinical

73 Trial Preparations

#### CONTROLLED TERM:

IN-VIVO \*FT; MOUSE \*FT; S.C. \*FT; SPLEEN-CELL \*FT;
PROLIFERATION \*FT; CELL-MEDIATED \*FT; IMMUNE-RESPONSE
\*FT; INTERFERON-GAMMA \*FT; INTERLEUKIN-4 \*FT; LAB.ANIMAL
\*FT: INJECTION \*FT: LYMPHOCYTE \*FT: IMMUNITY \*FT:

\*FT; INJECTION \*FT; LYMPHOCYTE \*FT; IMMUNITY \*FT; LAB.ANIMAL \*FT; INJECTION \*FT; LYMPHOCYTE \*FT;

IMMUNITY \*FT

[01] KEYHOLE-LIMPET-HEMOCYANIN \*PH; KEYHOLELI \*RN; VACCINE \*FT;

CONJUGATE \*FT; IMMUNIZATION \*FT; PH \*FT; PH \*FT

[02] GLUCAN-BETA-1,3-D \*PH; ADJUVAX \*PH; GLUCAB13D \*RN;

ADJUVANT \*FT; DRUG-COMPARISON \*FT; ALPHA-BETA-TECHNOL. \*FT;

IMMUNOSTIMULANTS \*FT; PH \*FT

[03] ALETHINE-BETA \*PH; DR9501199 \*RN; ADJUVANT \*FT;

2 Fig. 4 Tab.

DRUG-COMPARISON \*FT; ADJUVANTS \*FT; CYTOSTATICS \*FT; PH \*FT;

PH \*FT

BETAFECTIN \*PH; BETAFECTI \*RN; ADJUVANT \*FT; DRUG-COMPARISON [04]

\*FT; ALPHA-BETA-TECHNOL. \*FT; PH \*FT

[05] CRL-1005 \*PH; DR9703386 \*RN; ADJUVANT \*FT; DRUG-COMPARISON

\*FT; ADJUVANTS \*FT; VAXCEL \*FT; TRIAL-PREP. \*FT;

IMMUNOSTIMULANTS \*FT; PH \*FT; PH \*FT

[06] DETOX \*PH; RIBI-IMMUNOCHEM. \*FT; DETOX \*RN; DETOX \*RN;

ADJUVANT \*FT; DRUG-COMPARISON \*FT; ADJUVANTS \*FT;

IMMUNOSTIMULANTS \*FT; PH \*FT; ADJUVANTS \*FT;

IMMUNOSTIMULANTS \*FT; PH \*FT

COLONY-STIMULATING-FACTOR-GM \*PH; CSF-GM \*RN; ADJUVANT \*FT; [07]

DRUG-COMPARISON \*FT; LEXIGEN \*FT; PH \*FT; PH \*FT

QUILLAJA-SAPONIN \*PH; QS-21 \*PH; QUILLAJSA \*RN; ADJUVANT \*FT; [80]

DRUG-COMPARISON \*FT; AQUILA \*FT; ADJUVANTS \*FT;

IMMUNOSTIMULANTS \*FT; ADJUVANTS \*FT;

IMMUNOSTIMULANTS \*FT; PH \*FT

[09] TITERMAX \*PH; TITERMAX \*RN; ADJUVANT \*FT; DRUG-COMPARISON

\*FT; ADJUVANTS \*FT; CYTRX \*FT; PH \*FT

FIELD AVAIL.: AB; LA; CT FILE SEGMENT: Literature

L188 ANSWER 21 OF 63 DRUGU COPYRIGHT 2006 THE THOMSON CORP on STN

ACCESSION NUMBER: 1999-34186 DRUGU T M

Fluconazole versus amphotericin B for the prevention of TITLE:

fungal infection in neutropenic patients with hematologic

malignancy.

AUTHOR: Takatsuka H; Takemoto Y; Okamoto T; Fujimori Y; Tamura S;

Wada H; Okada M; Kanamaru A; Kakishita E

CORPORATE SOURCE: Hyogo-Coll.Med.; Univ.Kinki

Hyogo; Osaka; Jap. LOCATION:

SOURCE: Drugs Exp. Clin. Res. (25, No. 4, 193-200/ 1999)

35 Ref. CODEN: DECRDP

Second Department of Internal Medicine, Hyogu College of AVAIL. OF DOC.:

ISSN: 0378-6501 Medicine, 1-1 Mukogawacho, Nishinomiya, Hyogo 663-8501,

Japan. (e-mail: hematol@hyo-med.ac.jp).

LANGUAGE: English DOCUMENT TYPE: Journal

ABSTRACT:

P.o. fluconazole (FLC, Pfizer) reduced fungal infection to a greater extent than amphotericin B (AMB, Bristol-Myers Squibb) in a study of 124 patients with hematological malignancy and neutropenia. Although the use of idarubicin HCl by patients in the FLC group led to a greater reduction in WBC and absolute neutrophil levels, the febrile/neutropenic ratio and the beta-D glucan level were lower in the AMB group. FLC reduced the isolation of Candida albicans and C. glabrata from throat swabs and sputum. Concomitant treatment included polymyxin B tablets (Pfizer), amphotericin B suspension, inhaled amphotericin B and ciprofloxacin (Bayer). FLC may be more effective than AMB for fungal prophylaxis in patients with hematological malignancy and neutropenia.

SECTION HEADING: T Therapeutics

M Microbiology

CLASSIF. CODE: 53 Infection

55 Fungicides

CONTROLLED TERM:

NEUTROPENIA \*OC; ACUTE \*OC; MYELOGENOUS \*OC; LEUKEMIA \*OC; LYMPHOBLASTIC \*OC; PRELEUKEMIA \*OC; NONHODGKIN \*OC; LYMPHOMA \*OC; CHRON. \*OC; MYELOID \*OC; ADULT \*OC; THYMOCYTE

\*OC; INFECTION, FUNGUS \*TR; LYMPHOPROLIFERATIVE-DISEASE \*OC; MARROW-DISEASE \*OC; POLYMYXIN-B \*RC; AMPHOTERICIN-B \*RC; CIPROFLOXACIN \*RC; CASES \*FT; IN-VIVO \*FT; FUNGICIDE \*FT;

NEUTROPHIL \*FT; GLUCAN-BETA-1,3-D \*FT; CANDIDA \*FT; ALBICANS \*FT; GLABRATA \*FT; P.O. \*FT; LEUKOCYTE \*FT;

PROPHYLAXIS \*FT; THROAT \*FT; SPUTUM \*FT; FECES \*FT; KRUSEI \*FT; TROPICALIS \*FT; LEUKOCYTE \*FT; FUNGUS \*FT; ORL \*FT

FLUCONAZOLE \*TR; FLUCONAZOLE \*PH; PFIZER \*FT; IDARUBICIN \*RC;
FLUCONAZO \*RN; FUNGICIDES \*FT; TR \*FT; PH \*FT

CAS REGISTRY NO.: 86386-73-4

[02] AMPHOTERICIN-B \*TR; AMPHOTERICIN-B \*PH; BRISTOL-SQUIBB \*FT;

AMPHOTERI \*RN; ANTIBIOTICS \*FT; FUNGICIDES \*FT; TR \*FT; PH

\*FT

CAS REGISTRY NO.: 1397-89-3
FIELD AVAIL.: AB; LA; CT
FILE SEGMENT: Literature

L188 ANSWER 22 OF 63 DRUGU COPYRIGHT 2006 THE THOMSON CORP on STN

ACCESSION NUMBER: 1997-00016 DRUGU M T

TITLE: The use of glucan as immunostimulant in the treatment of

paracoccidioidomycosis.

AUTHOR: Meira D A; Pereira P C M; Marcondes Machado J; Mendes R P;

Barraviera B; Pellegrino J Jr; Rezkallah Iwasso M T; Peracoli M T S; Castilho L M; Thomazini I; Da Silva C L; Foss N T;

Curi P R

CORPORATE SOURCE: Univ.Sao-Paulo

LOCATION: Sao Paulo, Braz.

SOURCE: Am.J.Trop.Med.Hyg. (55, No. 5, 496-503, 1996) 5 Fig. 3 Tab.

45 Ref.

CODEN: AJTHAB ISSN: 0002-9637

AVAIL. OF DOC.: Departamento de Doencas Tropicais e Diagnostico por Imagem,

Faculdade de Medicina de Botucatu, Botucatu, Sao Paulo,

18618-000, Brazil.

LANGUAGE: English DOCUMENT TYPE: Journal

#### ABSTRACT:

[01]

Long-term i.v. beta-1,3D-glucan (GN) immunostimulation afforded an improved response to specific antifungal therapy in 10 male patients with predominantly severe paracoccidioidomycosis (Paracoccidioides brasiliensis) when compared to antifungal therapy on its own in 8 male patients with moderate paracoccidioidomycosis. After GN treatment, ESR values were higher, serum antibodies to P. brasiliensis were lower, the PHA skin-test gave a positive reaction and the helper-cell count tended to be higher. Moreover, during GN treatment, there was a rise in the serum tumor necrosis factor (TNF) level. Antifungal medication included amphotericin B (AB), ketoconazole (KC), sulfanilamide (SA), sulfadiazine (SD) and sulfamethoxazole plus trimethoprim (SM + TM).

SECTION HEADING: M Microbiology

T Therapeutics

CLASSIF. CODE: 20 Immunological

50 Biological Response Modifiers

53 Infection

CONTROLLED TERM:

GLUCAN-BETA-1,3-D \*TR; SEVERE \*TR; INFECTION, FUNGUS [01]

\*TR; AMPHOTERICIN-B \*RC; KETOCONAZOLE \*RC; GLUCAB13D \*RN; CASES \*FT; IN-VIVO \*FT; PARACOCCIDIOIDES \*FT; BRASILIENSIS

\*FT; LONG-TERM-THERAPY \*FT; I.V. \*FT; IMMUNOSTIMULANT \*FT; CELL-MEDIATED \*FT; IMMUNITY \*FT; HELPER-CELL \*FT;

COUNT \*FT; BLOOD-SERUM \*FT; TUMOR-NECROSIS-FACTOR \*FT; CONC.

\*FT; FUNGUS \*FT; INJECTION \*FT; LYMPHOCYTE \*FT;

THYMOCYTE \*FT; TR \*FT

FIELD AVAIL.:

AB; LA; CT

Literature FILE SEGMENT:

L188 ANSWER 23 OF 63 DRUGU COPYRIGHT 2006 THE THOMSON CORP on STN

ACCESSION NUMBER: 1996-11613 DRUGU

Pulmonary metastases neutralization and tumor rejection by in TITLE:

vivo administration of beta glucan and bispecific antibody.

AUTHOR:

Penna C; Dean P A; Nelson H

LOCATION:

Rochester, Minn., USA

SOURCE:

Int.J.Cancer (65, No. 3, 377-82, 1995) 5 Fig. 3 Tab. 21 Ref.

CODEN: IJCNAW ISSN: 0020-7136

AVAIL. OF DOC.:

Mayo Clinic, 200 First Street S.W. Rochester, MN 55905, USA.

(H.N.).

LANGUAGE:

English Journal

DOCUMENT TYPE:

## ABSTRACT:

The Authors tested whether i.p. beta glucan (Sigma-Chemical) could in situ-activate T cells, which could secondarily be retargeted with Dispecific antibodies (BsAbs; i.v.) to lyse tumor cells. The resulting anti-tumor effects were evaluated in a murine melanoma model. Therapeutic effects were then compared to those obtained with adoptively transferred, in vitro-activated lymphocytes retargeted with BsAb. In the neutralization model, there was a reduction in the number of metastases in the beta glucan + BsAb group vs. controls, and with beta glucan alone. In the established tumor model, beta glucan + BsAb reduced the incidence of s.c. tumors as compared with control, with BsAb alone and with beta glucan alone. It also prolonged survival of tumor-bearing mice compared with control, BsAb alone and beta glucan alone. T-cells can be activated by beta glucan and retargeted with F(ab')2 BsAb.

SECTION HEADING: P Pharmacology

CLASSIF. CODE:

20 Immunological

52 Chemotherapy - non-clinical

CONTROLLED TERM:

ANIMAL-NEOPLASM \*OC; LUNG \*OC; MELANOMA \*OC; METASTASIS \*OC; PNEUMOPATHY \*OC; CYTOSTATIC \*FT; IN-VITRO \*FT; IN-VIVO \*FT; INJECTION \*FT; LAB.ANIMAL \*FT; LYMPHOCYTE \*FT; MOUSE \*FT;

NEUTRALIZATION \*FT; PH \*FT; SURVIVAL \*FT; THYMOCYTE

[01] ANTIBODY \*FT; I.V. \*FT; MONOCLONAL \*FT

[02] GLUCAN-BETA-1,3-D \*PH; GLUCAB13D \*RN; SIGMA-CHEM.

\*FT; I.P. \*FT

CAS REGISTRY NO .: 9051-97-2 FIELD AVAIL.: AB; LA; CT

FILE SEGMENT: Literature

L188 ANSWER 24 OF 63 DRUGU COPYRIGHT 2006 THE THOMSON CORP on STN ACCESSION NUMBER: 1993-47257 DRUGU M S

Jones 10/630143

TITLE: Adverse Effects of Pefloxacin in Irradiated C3H/HeN Mice:

Correction with Glucan Therapy.

AUTHOR: Patchen M L; Brook I; Elliott T B; Jackson W E

LOCATION: Bethesda, Maryland, United States

SOURCE: Antimicrob.Agents Chemother. (37, No. 9, 1882-89, 1993) 4

Fig. 34 Ref.

CODEN: AMACCO ISSN: 0066-4804

AVAIL. OF DOC.: Department of Experimental Hematology, Armed Forces

Radiobiology Research Institute, Bethesda, Maryland

20889-5603, U.S.A.

LANGUAGE: English DOCUMENT TYPE: Journal

#### ABSTRACT:

Pefloxacin methanesulfonate dihydrate (PF, Rhone-Poulenc) given p.o. enhanced mortality following exposure of mice to radiation, despitereduction of translocation of bacteria from bloodstream to liver. I.v. glucan-beta-1,3-D (GC, from Sacch. cerevisiae) alone enhanced survival while GC + PF enhanced survival beyond that observed with GC alone. PF suppressed granulocyte-macrophage progenitor cell (GM-CFC) recovery; GC stimulated GM-CFC recovery while GC administered in combination with PF could override the hemopoietic suppressive effect of PF.

SECTION HEADING: M Microbiology

S Adverse Effects

CLASSIF. CODE: 20 Immunological

34 Toxicology

50 Biological Response Modifiers

54 Antiseptics

CONTROLLED TERM:

IN-VIVO \*FT; MOUSE \*FT; IRRADIATION \*FT; ALONE \*FT; COMB.
\*FT; MORTALITY \*FT; SURVIVAL \*FT; GRANULOCYTE \*FT; MACROPHAGE
\*FT; SPLEEN-CELL \*FT; LAB.ANIMAL \*FT; LEUKOCYTE \*FT; RES \*FT;

LYMPHOCYTE \*FT

[01] PEFLOXACIN \*PH; PEFLOXACIN \*AE; RHONE-POULENC \*FT;

INFECTION, BACT. \*OC; MARROW-DEPRESSION \*AE; MARROW-DISEASE \*AE; MESILATE \*PH; MESILATE \*AE; PEFLOXACI \*RN; P.O. \*FT; TOX. \*FT; ANTISEPTIC \*FT; ANTISEPTICS \*FT; PH \*FT; AE \*FT

CAS REGISTRY NO.: 70458-92-3

[02] GLUCAN-BETA-1,3-D \*PH; MARROW-DEPRESSION \*OC; MARROW-DISEASE \*OC; GLUCAB13D \*RN; I.V. \*FT; IMMUNOSTIMULANT \*FT; INJECTION \*FT; PH \*FT

CAS REGISTRY NO.: 9051-97-2
FIELD AVAIL.: AB; LA; CT
FILE SEGMENT: Literature

L188 ANSWER 25 OF 63 DRUGU COPYRIGHT 2006 THE THOMSON CORP on STN

ACCESSION NUMBER: 1991-44207 DRUGU C P

TITLE: Mitogenic and Colony-Stimulating Factor-Inducing Activities

of Polysaccharide Fractions from the Fruit Bodies of

Dictyophora indusiata FISCH.

AUTHOR: Hara C; Kumazawa Y; Inagaki K; Kaneko M; Kiho T; Ukai S

LOCATION: Gifu, Tokyo, Japan

SOURCE: Chem. Pharm. Bull. (39, No. 6, 1615-16, (1991)) 2 Tab. 18 Ref.

CODEN: CPBTAL ISSN: 0009-2363

AVAIL. OF DOC.: Shotoku Gakuen Women's Junior College, 1-38, Nakauzura, Gifu

500, Japan.

LANGUAGE: English DOCUMENT TYPE: Journal

#### ABSTRACT:

Mitogenic and colony stimulating factor (CSF) activities of 5 homogeneous polysaccharide and a conjugated polysaccharide fraction isolated from the fruiting bodies of Dictyophora indusiata were investigated. Fucomannogalactan (T-3-Ad) and conjugated polysaccharide fraction (T-2-A) showed significant mitogenic and CSF inducing activities (i.p. in mice). Of 2 beta-(1-6)-branched (1-3)-beta-D-glucans (T-4-N and T-5-N), only T-4-N showed both mitogenic and CSF inducing activities. Partially acetylated (1-3)-alpha-mannans (T-2-HN and T-3-M') were not significantly active.

SECTION HEADING: C Chemistry

P Pharmacology

CLASSIF. CODE: 20 Immunological

71 Medicinal Chemistry

CONTROLLED TERM:

ISOL. \*FT; DICTYOPHORA \*FT; INDUSIATA \*FT; FUNGUS \*FT;
MITOGEN \*FT; COLONY-STIMULATING-FACTOR \*FT; INDUCTION \*FT;

IMMUNOSTIMULANT \*FT; SPLEEN-CELL \*FT; IN-VITRO \*FT;
IN-VIVO \*FT; I.P. \*FT; MOUSE \*FT; LYMPHOCYTE \*FT;

INJECTION \*FT; LAB.ANIMAL \*FT

[01] GLUCAN-BETA-1,3-D \*OC; GLUCAN-BETA-1,3-D

\*PH; GLUCAB13D \*RN; OC \*FT; PH \*FT

CAS REGISTRY NO.: 9051-97-2

[02] MANNAN \*OC; MANNAN \*PH; MANNAN \*RN; OC \*FT; PH \*FT

CAS REGISTRY NO.: 51395-96-1

[03] POLYSACCHARIDE \*FT; OC \*FT; PH \*FT

FIELD AVAIL.: AB; LA; CT; MPC

FILE SEGMENT: Literature

L188 ANSWER 26 OF 63 DRUGU COPYRIGHT 2006 THE THOMSON CORP on STN

ACCESSION NUMBER: 1991-37445 DRUGU P

TITLE: Oral Administration of SSG, a beta-Glucan Obtained from

Sclerotinia sclerotiorum, Affects the Function of Peyer's

Patch Cells.

AUTHOR: Hashimoto K; Suzuki I; Yadomae T

LOCATION: Tokyo, Japan

SOURCE: Int.J.Immunopharmacol. (13, No. 4, 437-42, 1991) 1 Fig. 1

Tab. 26 Ref.

CODEN: IJIMDS ISSN: 0192-0561

AVAIL. OF DOC.: Laboratory of Immunopharmacology of Microbial Products, Tokyo

College of Pharmacy, Horinouchi 1432-1, Hachioji, Tokyo

192-03, Japan.

LANGUAGE: English DOCUMENT TYPE: Journal

# ABSTRACT:

The effect of orally administered SSG, a beta-1,3-glucan obtained from the culture filtrate of the fungus Sclerotinia sclerotiorum IFO 9395, on the function of Peyer's patch (PP) cells was investigated in comparison with that on spleen cells in mice. Oral SSG enhanced the proliferative response of PP cells to a T-mitogen, concanavalin A (Con A), and a B cell mitogen, lipopolysaccharide (LPS), although the response of spleen cells was unaffected. PP cells taken from mice exposed previously to oral SSG showed enhanced

plaque-forming cell responses to sheep red blood cells (SRBC) after antigen (SRBC) stimulation for 5 days in-vitro. Results show that SSG can modulate the mucosal immune response.

SECTION HEADING: P Pharmacology

CLASSIF. CODE: 16 Gastrointestinal

20 Immunological

50 Biological Response Modifiers

CONTROLLED TERM:

[01] GLUCAN-BETA-1,3-D \*PH; CONCANAVALIN-A \*RC;

GLYCOLIPID \*RC; P.O. \*FT; MOUSE \*FT; IN-VIVO \*FT; IMMUNOSTIMULANT \*FT; MUCOSA \*FT; PEYER-PATCH \*FT; SPLEEN-CELL \*FT; PROLIFERATION \*FT; IMMUNE-RESPONSE \*FT; GASTROINTEST. \*FT; THYMOCYTE \*FT; INTESTINE \*FT; LAB.ANIMAL \*FT; LYMPHOCYTE \*FT; IMMUNITY \*FT;

GLUCAB13D \*RN; PH \*FT

CAS REGISTRY NO.: 9051-97-2
FIELD AVAIL.: AB; LA; CT
FILE SEGMENT: Literature

L188 ANSWER 27 OF 63 DRUGU COPYRIGHT 2006 THE THOMSON CORP on STN

ACCESSION NUMBER: 1990-47819 DRUGU F

TITLE: Enhancement of Hematopoietic Response of Mice by

Intraperitoneal Administration of a beta-Glucan, SSG,

Obtained from Sclerotinia sclerotiorum.

AUTHOR: Hashimoto K; Suzuki I; Ohsawa M; Oikawa S; Yadomae T

CORPORATE SOURCE: Nippon-Beet-Sugar

LOCATION: Tokyo, Japan

SOURCE: J. Pharmacobiodyn. (13, No. 8, 512-17, 1990) 4 Fig. 1 Tab. 20

Ref.

CODEN: JOPHDQ ISSN: 0386-846X

AVAIL. OF DOC.: Tokyo College of Pharmacy, Horinouchi, Hachioji, Tokyo

192-03, Japan.

LANGUAGE: English DOCUMENT TYPE: Journal

ABSTRACT:

In mice, i.p. SSG (a beta-glucan isolated from Sclerotinia sclerotiorum) markedly increased the % of PMN in both spleen and peripheral blood, increased the numbers of macrophage progenitor cells in both spleens and femurs, and increased levels of colony-stimulating activity (CSA) in sera. The results suggest that i.p. SSG in mice enhances the production of colony-stimulating factors (CSFs) and then increases the numbers of both spleen and peripheral blood leukocytes.

SECTION HEADING: P Pharmacology

CLASSIF. CODE: 20 Immunological

50 Biological Response Modifiers 52 Chemotherapy - non-clinical

CONTROLLED TERM:

[01] GLUCAN-BETA-1,3-D \*PH; IN-VIVO \*FT; MOUSE \*FT; I.P.

\*FT; MARROW \*FT; POLYMORPHONUCLEAR \*FT; SPLEEN \*FT; BLOOD

\*FT; MACROPHAGE \*FT; PERIPHERAL \*FT; MONOCYTE \*FT;

BLOOD-SERUM \*FT; LYMPHOCYTE \*FT;

COLONY-FORMING-UNIT \*FT; PERITONEAL \*FT; LAB.ANIMAL \*FT;

Searched by Barb O'Bryen, STIC 2-2518

INJECTION \*FT; LEUKOCYTE \*FT; RES \*FT; GLUCAB13D \*RN; PH \*FT

FIELD AVAIL.: AB; LA; CT FILE SEGMENT: Literature

L188 ANSWER 28 OF 63 DRUGU COPYRIGHT 2006 THE THOMSON CORP on STN

ACCESSION NUMBER: 1989-21277 DRUGU C P

TITLE: Antitumor and Immunomodulating Activities of a beta-Glucan

Obtained from Liquid-Cultured Grifola Frondosa.

AUTHOR: Suzuki I; Hashimoto K; Oikawa S; Sato K; Osawa M; Yadomae T

CORPORATE SOURCE: Nippon-Beet-Sugar

LOCATION: Tokyo, Japan

SOURCE: Chem. Pharm. Bull. (37, No. 2, 410-13, 1989) 1 Fig. 4 Tab. 16

Ref.

CODEN: CPBTAL ISSN: 0009-2363

AVAIL. OF DOC.: Tokyo College of Pharmacy, Horinouchi, Hachioji, Tokyo

192-03, Japan.

LANGUAGE: English DOCUMENT TYPE: Journal

ABSTRACT:

A beta-1,3-glucan (LELFD) from Grifola frondosa mycelium had i.p. and intra-lesional (i.l.) antitumor activity against Meth A fibrosarcoma and IMC carcinoma in mice but, like lentinan (LE, Yamanouchi), was without effect on L1210 and P388 leukemias. The activities of natural killer (NK) spleen cells and macrophages in mice were enchanced by i.p. LELFD which also potentiated antibody responses to sheep RBC and activated the alternate complement pathway.

SECTION HEADING: C Chemistry

P Pharmacology

CLASSIF. CODE: 20

20 Immunological

50 Biological Response Modifiers 52 Chemotherapy - non-clinical

71 Medicinal Chemistry

CONTROLLED TERM:

[01] GLUCAN-BETA-1,3-D \*OC; GLUCAN-BETA-1,3-D

\*PH; P388 \*OC; ANIMAL-NEOPLASM \*OC; LEUKEMIA \*OC; L1210 \*OC; METH-A \*OC; FIBROSARCOMA \*OC; CARCINOMA \*OC; LENTINAN \*RC; PICIBANIL \*RC; ISOL. \*FT; GRIFOLA \*FT; FRONDOSA \*FT; IN-VITRO \*FT; IN-VIVO \*FT; I.P. \*FT; INTRATUMOR \*FT; CYTOSTATIC \*FT;

MOUSE \*FT; IMMUNOSTIMULANT \*FT; NAT.KILLER-CELL

\*FT; STIMULATION \*FT; MACROPHAGE \*FT; ACTIVATION \*FT; YAMANOUCHI \*FT; COMPLEMENT \*FT; ANTIBODY-RESPONSE \*FT; MUSHROOM \*FT; INJECTION \*FT; LAB.ANIMAL \*FT; LYMPHOCYTE \*FT; RES \*FT; IMMUNITY \*FT; GLUCAB13D \*RN; OC \*FT; PH

\*FT

FIELD AVAIL .: AB; LA; CT; MPC

FILE SEGMENT: Literature

L188 ANSWER 29 OF 63 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 2

ACCESSION NUMBER: 2001:635920 CAPLUS

DOCUMENT NUMBER: 135:185438

TITLE: Non-antigenic mucosal adjuvant formulation

INVENTOR(S): Raa, Jan; Berstad, Aud Kathrine Herland; Bakke, Hilde;

Haneberg, Bjorn; Haugen, Inger Lise; Holst, Johan;

Janakova, Liba; Korsvold, Gro Ellen; Oftung, Fredrik Biotec Asa, Norway

PATENT ASSIGNEE(S): SOURCE:

PCT Int. Appl., 19 pp

CODEN: PIXXD2

OCCUMENT TYPE: Patent

DOCUMENT TYPE:

English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

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KIND DATE
                                         APPLICATION NO. DATE
    PATENT NO.
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    _____
    WO 2001062283
                        A2
                               20010830
                                        WO 2001-IB144
                                                                20010202
                        A3
                               20020214
    WO 2001062283
        W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
            CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR,
            HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT,
            LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU,
            SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN,
            YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
        RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
            DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
            BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
                         A1
                               20020124
                                        US 2000-511582
                                                                  20000223
    US 2002009463
                                         EP 2001-912024
    EP 1259259
                         A2
                               20021127
                                                                  20010202
        R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
            IE, SI, LT, LV, FI, RO, MK, CY, AL, TR
                                        JP 2001-561347
                               20030805
                                                                  20010202
    JP 2003523401
                        T2
    AU 771205
                        B2
                                          AU 2001-40943
                                                                 20010202
                               20040318
                               20021015 NO 2002-3935
    NO 2002003935
                        Α
                                                                 20020819
                        A1
                                          US 2002-203280
    US 2003104010
                                                                 20021007
                               20030605
                                                             A 20000223
W 20010202
PRIORITY APPLN. INFO.:
                                           US 2000-511582
                                           WO 2001-IB144
    Entered STN: 31 Aug 2001
ED
    An adjuvant for mucosal vaccines which modulates the effects of
AB
    substances, including vaccine antigens in contact with mucosal body
    surfaces is described. Thus, a formulation containing (1\rightarrow 3)-\beta-D-
    glucan enhanced the prodn.of IgG against influenza vaccine antigens.
IC
    ICM A61K039-39
         A61K039-00; A61K039-145; A61K031-573; A61P037-04; A61P037-08;
         A61P011-08; A61P019-02; A61P031-16
    63-3 (Pharmaceuticals)
CC
IT
    Immunostimulants
        (adjuvants; non-antigenic mucosal adjuvant vaccine formulation)
IT
    Allergy
    Arthritis
    Spleen
      T cell (lymphocyte)
    Vaccines
        (non-antigenic mucosal adjuvant vaccine formulation)
    9051-97-2, \beta-D- Glucan, (1\rightarrow 3)-
    37361-00-5, \beta-D- Glucan, (1\rightarrow 6)-
    RL: BAC (Biological activity or effector, except adverse); BSU
     (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (non-antigenic mucosal adjuvant vaccine formulation)
```

L188 ANSWER 30 OF 63 CAPLUS COPYRIGHT 2006 ACS on STN DUPLICATE 3

ACCESSION NUMBER: 2001:482903 CAPLUS

DOCUMENT NUMBER: 135:282915

TITLE: Th1/Th2-balancing immunomodulating activity of

gel-forming  $(1\rightarrow 3)$ - $\beta$ -glucans from fungi

AUTHOR(S): Suzuki, Yoko; Adachi, Yoshiyuki; Ohno, Naohito;

Yadomae, Toshiro

CORPORATE SOURCE: Laboratory of Immunopharmacology of Microbial

Products, School of Pharmacy, Tokyo University of Pharmacy and Life Science, Tokyo, 192-0392, Japan Biological & Pharmaceutical Bulletin (2001) 24(7)

SOURCE: Biological & Pharmaceutical Bulletin (2001), 24(7),

811-819

CODEN: BPBLEO; ISSN: 0918-6158
Pharmaceutical Society of Japan

PUBLISHER: Pharmaceutical Society of Japan

DOCUMENT TYPE: Journal LANGUAGE: English ED Entered STN: 05 Jul 2001

The immunomodulating effects of various gel-forming  $(1\rightarrow 3)-\beta$ -AΒ glucans, grifolan (GRN), SSG, sonifilan (SPG) and alkaline-treated SPG (SPG-OH), on balancing helper T cell activity were examined in a murine model. Plasma from mice that were injected with GRN or SPG-OH and trinitrophenyl ovalbumin (TNP-OVA) contained TNP-specific antibodies of both IgG1 and IgG2a isotypes. Administration of SSG and TNP-OVA significantly augmented the synthesis of IgG2a antibodies, while the synthesis of IgG1 was reduced. However, SPG did not enhance the antibody response. In the culture supernatants of splenocytes obtained from GRNor SPG-OH-administered mice, high levels of IgG1 and low levels of IgG2a and IFN  $\gamma$  were detected. In contrast, high levels of IgG2a and IFN  $\gamma$  and low levels of IqG1 were detected in the case of administration of SSG. Furthermore, it was shown by intracellular cytokine staining that the proportion of IFN  $\gamma$ +CD4+ double-pos. cells among the CD4+ cells from mice administered SSG was most strongly increased by addition of PMA and A23187. On the other hand, the expression of IL-12 p40 mRNA was more markedly elevated in splenocytes after combined administration of TNP-OVA plus SSG than after administration of TNP-OVA alone. The highest IFN γ production was observed when adherent cells of mice administered TNP-OVA and SSG were cultured with TNP-primed lymphocytes. This effect of administration of SSG on IFN- $\gamma$  production was completely inhibited by addition of anti-IL-12 mAb. In conclusion, our study showed that β-glucans have various effects on the Th1 or Th2-dependent antibody subclasses, in particular, SSG induces the development of Th1 cells via the IL-12 pathway.

CC 1-7 (Pharmacology)

IT Polysaccharides, biological studies

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(Th1/Th2-balancing immunomodulating activity of gel-forming  $(1\rightarrow 3)$ - $\beta$ - glucans from fungi)

IT Immunostimulants

(adjuvants; Th1/Th2-balancing immunomodulating activity of gel-forming  $(1\rightarrow 3)-\beta$ -glucans from fungi)

IT T cell (lymphocyte)

(helper cell/inducer, TH1; Th1/Th2-balancing immunomodulating activity of gel-forming  $(1\rightarrow 3)-\beta$ -glucans from fungi)

IT T cell (lymphocyte)

(helper cell/inducer, TH2; Th1/Th2-balancing immunomodulating activity of gel-forming  $(1\rightarrow 3)$ - $\beta$ -glucans from fungi)

IT 9050-67-3, Sonifilan 9050-67-3D, Sonifilan, single helical conformer
9051-97-2, SSG 104074-36-4, Grifolan

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(Th1/Th2-balancing immunomodulating activity of gel-forming

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Jones
                                        10/630143
                 glucans from fungi)
        (1→3)-β-
REFERENCE COUNT:
                        46
                              THERE ARE 46 CITED REFERENCES AVAILABLE FOR THIS
                              RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
L188 ANSWER 31 OF 63 CAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER:
                        2006:32968 CAPLUS
                        Tinospora polysaccharide as immunostimulant for
TITLE:
                        treatment of proliferative diseases or infection
                        Nair, P. k. Raveendran; Melnick, Steven J.;
INVENTOR(S):
                        Ramachandran, Cheppail
PATENT ASSIGNEE(S):
                        USA
SOURCE:
                        U.S. Pat. Appl. Publ., 43 pp.
                        CODEN: USXXCO
DOCUMENT TYPE:
                        Patent
                        English
LANGUAGE:
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
     KIND
                                      APPLICATION NO. DATE
    PATENT NO.
                               DATE
                               -----
                                                                20050711
    US 2006009501
                               20060112 US 2005-178620
                        A1
                               20060126
                                        WO 2005-US24410
    WO 2006010069
                        A1
        W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
            CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
            GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ,
            LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA,
            NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK,
            SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU,
            ZA, ZM, ZW
        RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE,
            IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ,
            CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH,
            GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
            KG, KZ, MD, RU, TJ, TM
PRIORITY APPLN. INFO.:
                                           US 2004-586548P P 20040709
    Entered STN: 13 Jan 2006
ED
    The invention concerns a novel polysaccharide. RR1 is an \alpha-D-glucan
AB
    polysaccharide composed of a (1-4) linked back bone and
     (1-6) linked branches, which has been isolated from a medicinal
    herb, Tinospora cordifolia. RR1 exhibits unique immune-stimulating
    properties, is non-cytotoxic, and non-proliferating to normal lymphocytes,
    as well as tumor cell lines. The subject invention also concerns compns.
    containing an RR1 compound and methods for modulating an immune response in a
    subject using RR1 compds. The invention also provides methods for the use
    of an RR1 compound in conjunction with an antigen to stimulate an immune
    response, the RR1 compound providing an adjuvant-like activity in the
    generation of a Th1-type immune response to the antigen.
INCL 514367000
    1-7 (Pharmacology)
CC
    Section cross-reference(s): 11, 15
    INDEXING IN PROGRESS
IT
IT
    Polysaccharides
    RL: NPO (Natural product occurrence); PAC (Pharmacological
```

Searched by Barb O'Bryen, STIC 2-2518

(1,6)-α-D-qlycosidic linked side chain; Tinospora polysaccharide as immunostimulant for treatment of proliferative diseases or

activity); PRP (Properties); PUR (Purification or recovery); THU (Therapeutic use); BIOL (Biological study); OCCU (Occurrence); PREP

 $(D-glucan; (1,4)-\alpha-D-glycosidic linkage in main chain$ 

(Preparation); USES (Uses)

infection)

```
IT
    Cell activation
        (T cell; Tinospora polysaccharide as immunostimulant for
        treatment of proliferative diseases or infection)
IT
    Anti-infective agents
     Antitumor agents
     Combination chemotherapy
    Drugs
     Embryophyta
     Human
       Immunostimulants
     Immunotherapy
     Infection
    Leukemia
    Macrophage
     Phagocytosis
     Tinospora cordifolia
        (Tinospora polysaccharide as immunostimulant for treatment of
        proliferative diseases or infection)
IT
    B cell (lymphocyte)
    Macrophage
       T cell (lymphocyte)
     Transcriptional regulation
        (activation; Tinospora polysaccharide as immunostimulant for treatment
        of proliferative diseases or infection)
IT
     Immunostimulants
        (adjuvants; Tinospora polysaccharide as immunostimulant for treatment
        of proliferative diseases or infection)
IT
     Growth factors, animal
     Interferons
     Interleukins
     Lymphokines
     Platelet-derived growth factors
     Tumor necrosis factors
     RL: PAC (Pharmacological activity); THU (Therapeutic
     use); BIOL (Biological study); USES (Uses)
        (and \alpha\text{-D-} glucan polysaccharide; pharmaceutical compns.
        containing; Tinospora polysaccharide as immunostimulant for
        treatment of proliferative diseases or infection)
     T cell (lymphocyte)
IT
        (helper cell/inducer, TH1, -stimulated cytokine production; Tinospora
        polysaccharide as immunostimulant for treatment of proliferative
        diseases or infection)
     9074-78-6DP, \alpha-D- Glucan, branched
IT
     RL: NPO (Natural product occurrence); PAC (Pharmacological
     activity); PRP (Properties); PUR (Purification or recovery); THU
     (Therapeutic use); BIOL (Biological study); OCCU (Occurrence); PREP
     (Preparation); USES (Uses)
        (RR1; Tinospora polysaccharide as immunostimulant for
        treatment of proliferative diseases or infection)
TT
     9008-22-4, Laminarin
     RL: BSU (Biological study, unclassified); PAC (Pharmacological
     activity); BIOL (Biological study)
        (Tinospora polysaccharide as immunostimulant for treatment of
        proliferative diseases or infection)
     9061-61-4, Nerve growth factor
                                      139639-23-9, tissue plasminogen activator
IT
     RL: PAC (Pharmacological activity); THU (Therapeutic
     use); BIOL (Biological study); USES (Uses)
        (and \alpha-D- glucan polysaccharide; pharmaceutical compns.
        containing; Tinospora polysaccharide as immunostimulant for
        treatment of proliferative diseases or infection)
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Jones 10/630143
L188 ANSWER 32 OF 63 CAPLUS COPYRIGHT 2006 ACS on STN
                          2004:902155 CAPLUS
ACCESSION NUMBER:
                          141:384286
DOCUMENT NUMBER:
TITLE:
                          Novel encochleation methods, cochleates and methods of
                          Mannino, Raphael J.; Gould-Fogerite, Susan;
INVENTOR (S):
                          Krause-Elsmore, Sara L.; Delmarre, David; Lu, Ruying
                          Biodelivery Sciences International, Inc., USA;
PATENT ASSIGNEE(S):
                          University of Medicine and Dentistry of New Jersey
                          PCT Int. Appl., 195 pp.
SOURCE:
                          CODEN: PIXXD2
DOCUMENT TYPE:
                          Patent
                          English
LANGUAGE:
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
     PATENT NO.
                     KIND DATE APPLICATION NO.
                                                                     DATE
                                 -----
                         ----
                                             WO 2004091578
                                                                      20040409
                         A2 20041028 WO 2004-US11026
                          C1
                                20050127
     WO 2004091578
     WO 2004091578
                          A3 20050331
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
             CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
             GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KA, KZ, LC,
             LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI,
             NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
             TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
         RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ,
             BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE,
             ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI,
             SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN,
             TD, TG
     US 2005013854
                          A1 20050120
                                              US 2004-822230 20040409

US 2003-461483P P 20030409

US 2003-463076P P 20030415

US 2003-499247P P 20030828

US 2003-502557P P 20030911

US 2003-532755P P 20031224

US 2004-537252P P 20040115

US 2004-556192P P 20040324
                                              US 2004-822230
                                                                       20040409
PRIORITY APPLN. INFO.:
     Entered STN: 28 Oct 2004
ED
     The invention generally relates to cochleate drug delivery vehicles.
AΒ
```

- AB The invention generally relates to cochleate drug delivery vehicles. Disclose are novel methods for making cochleates and cochleate compns. that include introducing a cargo moiety to a liposome in the presence of a solvent. Also disclosed are cochleates and cochleate compns. that include an aggregation inhibitor, and optionally, a cargo moiety. Addnl., anhydrous cochleates that include a protonized cargo moiety, a divalent metal cation and a neg. charge lipid are disclosed. Methods of using the cochleate compns. of the invention, including methods of administration, are also disclosed.
- IC ICM A61K009-127
- CC 63-6 (Pharmaceuticals)
  - Section cross-reference(s): 1, 2, 17, 18
- IT Blood coagulation

# Immunity

(disorder; novel encochleation methods and cochleates and methods of use for delivery of drugs and other agents using liposomes and aggregation inhibitors)

IT Adenoma

Aggregation Alopecia Alzheimer's disease Analgesics Anesthetics Animal virus Anti-Alzheimer's agents Anti-infective agents Antiarthritics Antiasthmatics Antibacterial agents Antibiotics Anticholesteremic agents Anticoagulants Anticonvulsants Antidepressants Antidiabetic agents Antihistamines Antihypertensives Antihypotensives Antimicrobial agents Antiobesity agents Antioxidants Antiparkinsonian agents Antipsychotics Antirheumatic agents Antitumor agents Antiviral agents Arthritis Asthma Atherosclerosis Autoimmune disease Biliary tract, neoplasm Carcinoma Carcinoma Cations Chelating agents Cholinergic antagonists Cognition enhancers Cystic fibrosis Cytoprotective agents Cytotoxic agents Dairy products Decongestants Detergents Eczema Esophagus, neoplasm Expectorants Flavoring materials Fungicides Gene therapy Genetic vectors Ginkgo Gout Graves' disease Gums and Mucilages Headache Hemophilia. Hemostatics Hypercholesterolemia

Hyperglycemia Hypericum Hypertension Hypolipemic agents Hypotension Imaging agents Immunostimulants Immunosuppressants Infection Inflammation Leukemia Leukotriene antagonists Lung, neoplasm Lymphoma Malnutrition Mammary gland, neoplasm Melanoma Milk Mouthwashes Multiple sclerosis Muscular dystrophy Myasthenia gravis Mycosis Neoplasm Neuroglia, neoplasm Nutrients Obesity Organelle Osteoarthritis Ovary, neoplasm Packaging materials Pain Pancreas, neoplasm Parasiticides Parkinson's disease Pigments, biological Plasmids Prostate gland, neoplasm Psoriasis Psychotropics Rheumatoid arthritis Sarcoma Schizophrenia Skin, disease Stomach, neoplasm Sweetening agents Testis, neoplasm Tranquilizers Transplant rejection

# Uterus, neoplasm Vaccines

Vasoconstrictors

Vasodilators

TТ

(novel encochleation methods and cochleates and methods of use for delivery of drugs and other agents using liposomes and aggregation inhibitors)

1398-61-4, Chitin 4004-05-1, DOPE 9000-01-5, Acacia, gum 9000-07-1, Carrageenan 9000-65-1, Gums, tragacanth 9000-69-5, Pectin 9002-89-5, Polyvinyl alcohol 9003-01-4, Polyacrylic acid 9003-39-8, Polyvinylpyrrolidone 9004-32-4, Carboxymethyl cellulose 9004-42-6,

9004-53-9, Dextrin 9004-57-3, Ethylcellulose Carboxyethyl cellulose 9004-62-0, Hydroxyethyl cellulose 9004-64-2, Hydroxypropyl cellulose 9004-65-3, Hydroxypropylmethyl cellulose 9004-67-5, Methylcellulose 9005-18-9, Propylcellulose 9005-25-8D, Starch, hydroxypropylated 9005-82-7, Amylose high-amylose 9005-38-3, Sodium alginate 9012-76-4, Chitosan 9013-95-0, Levan 9057-02-7, Pullulan 14127-61-8, Calcium(2+), biological studies 22541-12-4, Barium(2+), biological 25322-68-3, Polyethylene glycol 37353-59-6, Hydroxymethyl 66457-06-5, Elsinan 70614-14-1, Dioleoylphosphatidylserine RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (novel encochleation methods and cochleates and methods of use for delivery of drugs and other agents using liposomes and aggregation inhibitors)

IT 50-99-7, Glucose, biological studies 57-50-1, Sucrose,
 biological studies 69-79-4, Maltose 81-07-2, Saccharine 9050-36-6,
 Maltodextrin 22839-47-0, Aspartame 64519-82-0, Isomalt
 RL: FFD (Food or feed use); BIOL (Biological study); USES (Uses)
 (sweetening agent; novel encochleation methods and cochleates and
 methods of use for delivery of drugs and other agents using liposomes
 and aggregation inhibitors)

IT 9012-72-0, Glucan

RL: BSU (Biological study, unclassified); BIOL (Biological study) (synthesis, inhibitors; novel encochleation methods and cochleates and methods of use for delivery of drugs and other agents using liposomes and aggregation inhibitors)

L188 ANSWER 33 OF 63 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2004:252369 CAPLUS

DOCUMENT NUMBER: 140:269531

TITLE: Autologous ghrelin and encoding nucleic acid and

for<u>eign T cell epitope c</u>onjugates for vaccination against obesity and excess body fat increase or loss

in human and animal

INVENTOR(S): Boving, Time Elisabeth Gottschalk; Klysner, Steen

PATENT ASSIGNEE(S): Pharmexa A/s, Den.
SOURCE: PCT Int. Appl., 83 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.					KIND DATE				APPL:	ICAT:	DATE						
						-											
WO 2004024183					A1		2004	0325	,	WO 2	003-1	20030912					
WO 2004024183				B1		2004	0513										
	W:	ΑE,	AG,	ΑĹ,	AM,	AT,	AU,	ΑZ,	BA,	BB,	BG,	BR,	BY,	ΒZ,	CA,	CH,	CN,
		CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DΖ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,	GE,
		GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KΕ,	KG,	KP,	KR,	ΚZ,	LC,	LK,
		LR,	LS,	LT,	LU,	LV,	MΑ,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NI,	NO,	NZ,
		OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	SY,	ΤJ,	TM,
		TN,	TR,	TT,	TZ,	UA,	UG,	US,	UΖ,	VC,	VN,	YU,	ZA,	ZM,	ZW		
	RW:	GH,	GM,	KE,	LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,	ΑZ,	BY,
		KG,	KZ,	MD,	RU,	ΤJ,	TM,	ΑT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	EE,	ES,
		FI,	FR,	GB,	GR,	HU,	IE,	IT,	LU,	MC,	NL,	PT,	RO,	SE,	SI,	SK,	TR,
		BF,	ВJ,	CF,	CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML,	MR,	ΝE,	SN,	TD,	TG
CA 2498739			AA 20040325				CA 20	003-	20	0030	912						
EP 1539232				A1	20050615				EP 20	003-	20030912						
	R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,
		IE,	SI,	LT,	LV,	FI,	RO,	MK,	CY,	AL,	TR,	BG,	CZ,	EE,	HU,	SK	

NO 2005001779 A 20050411 NO 2005-1779 20050411
PRIORITY APPLN. INFO:: DK 2002-1345 A 20020912
US 2002-410164P P 20020912
WO 2003-DK592 W 20030912

ED Entered STN: 26 Mar 2004

AB Disclosed are novel methods that generally rely on immunization against autologous ghrelin. Immunization is preferably effected by administration of analogs of autologous ghrelin, said analogs being capable of inducing antibody production against the autologous ghrelin polypeptides. Especially preferred as an immunogen is autologous ghrelin, which has been modified by introduction of one single or a few foreign, immunodominant and promiscuous T-cell epitopes. Also disclosed are nucleic acid vaccination against ghrelin and vaccination using live vaccines as well as methods and means useful for the vaccination. Such methods and means include methods for the preparation of analogs and pharmaceutical formulations, as well as nucleic acid fragments, vectors, transformed cells, polypeptides and pharmaceutical formulations.

IC ICM A61K039-39

ICS A61K039-385; A61K039-00; C07K014-435; A61P003-04

CC 15-2 (Immunochemistry)

Section cross-reference(s): 3, 63

IT Animal cell line

(S2; autologous ghrelin and encoding nucleic acid and foreign  ${\bf T}$  cell epitope conjugates for vaccination against obesity and excess body fat increase or loss)

IT Animal cell line

(SF9; autologous ghrelin and encoding nucleic acid and foreign **T** cell epitope conjugates for vaccination against obesity and excess body fat increase or loss)

IT Immunostimulants

(adjuvants; autologous ghrelin and encoding nucleic acid and foreign T cell epitope conjugates for vaccination against obesity and excess body fat increase or loss)

IT Immune tolerance

# Vaccines

(auto-; autologous ghrelin and encoding nucleic acid and foreign T cell epitope conjugates for vaccination against obesity and excess body fat increase or loss)

IT Amide group

Animal

Animal cell

Animal cell line

Anorexia

Antigen presentation

Antigen-presenting cell

Bos taurus

Burn

Cachexia

Canis familiaris

DNA sequences

**Epitopes** 

Eubacteria

Eukaryota

Fungi

Genetic vectors

Human

## Immunostimulants

Immunotherapy
Influenza virus
Microorganism

```
Molecular cloning
     Mus
     Obesity
     PCR (polymerase chain reaction)
     Plant cell
     Plasmodium falciparum
     Prokaryota
     Protein sequences
     Protozoa
     Rattus
     Sterculia urens
     Sus scrofa domestica
     Viral vectors
     brupW
     Yeast
     cDNA sequences
        (autologous ghrelin and encoding nucleic acid and foreign T
        cell epitope conjugates for vaccination against obesity and excess body
        fat increase or loss)
IT
     B cell (lymphocyte)
       T cell (lymphocyte)
        (epitope; autologous ghrelin and encoding nucleic acid and foreign
        T cell epitope conjugates for vaccination against obesity and
        excess body fat increase or loss)
IT
     T cell (lymphocyte)
        (helper cell, epitope; autologous ghrelin and encoding nucleic acid and
        foreign T cell epitope conjugates for vaccination against
        obesity and excess body fat increase or loss)
IT
     Animal cell
        (mammalian; autologous ghrelin and encoding nucleic acid and foreign
        T cell epitope conjugates for vaccination against obesity and
        excess body fat increase or loss)
IT
     541-59-3, Maleimide 1398-61-4, Chitin 7693-46-1, p-Nitrophenyl
                  8063-16-9, Psyllium 9000-01-5, Gum arabic 9000-07-1, 9000-21-9, Furcellaran 9000-28-6, Gum ghatti 9000-30-0
     chloroformate
     Carrageenan
                                                                    9000-30-0,
            9000-40-2, Locust bean gum 9000-65-1, Tragacanth 9000-69-5,
              9002-84-0, Polytetrafluoroethylene
                                                    9002-89-5, Poly(vinyl
     Pectin
     alcohol)
                9002-98-6, PEI 9003-01-4, Polyacrylic acid
                                                              9003-05-8,
                      9003-39-8, Poly(vinyl pyrrolidone))
     Polyacrylamide
                                                             9004-34-6,
     Cellulose, biological studies 9004-54-0, Dextran, biological studies
     9005-25-8, Starch, biological studies 9005-32-7D, Alginic acid, derivs.
     9005-79-2, Glycogen, biological studies 9011-14-7, Poly(methyl
     methacrylate)
                     9012-36-6, Agarose 9012-72-0, Glucan
     9012-76-4, Chitosan
                           9014-63-5, Xylan
                                              9036-88-8, Mannan
                                                                   9037-22-3,
                  9057-02-7, Pullulan
                                        11078-30-1, Galactomannan
     Amylopectin
                          12619-70-4D, Cyclodextrin, derivs.
     11138-66-2, Xanthan
                                                                 24937-78-8,
     Poly(ethylene-co-vinyl acetate)
                                       25087-26-7, Polymethacrylic acid
     25249-16-5, Poly(2-hydroxyethyl methacrylate) 25322-68-3D, Polyethylene
                       26780-50-7D, Poly(lactide-co-glycolide), derivs.
     glycol, derivs.
     37294-28-3, Xyloglucan 51751-43-0D, vinylene derivs.
                83869-56-1, GM-CSF
                                     110865-71-9, Acetan
     Tamarine
     RL: BSU (Biological study, unclassified); THU (Therapeutic use);
     BIOL (Biological study); USES (Uses)
        (autologous ghrelin and encoding nucleic acid and foreign T cell
        epitope conjugates for vaccination against obesity and excess body fat
        increase or loss)
REFERENCE COUNT:
                               THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS
                               RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
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L188 ANSWER 34 OF 63 CAPLUS COPYRIGHT 2006 ACS on STN

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2004:493559 CAPLUS
ACCESSION NUMBER:
                         141:37601
DOCUMENT NUMBER:
                         Therapy-enhancing glucan
TITLE:
                         Cheung, Nai-kong V.
INVENTOR(S):
PATENT ASSIGNEE(S):
                         Sloan-Kettering Institute for Cancer Research, USA
SOURCE:
                         U.S. Pat. Appl. Publ., 46 pp., Cont.-in-part of Appl.
                         No. PCT/US02/01276.
                         CODEN: USXXCO
DOCUMENT TYPE:
                         Patent
LANGUAGE:
                         English
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
     PATENT NO.
                       KIND
                               DATE
                                         APPLICATION NO.
                                                                  DATE
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                               20040617 US 2003-621027
     US 2004116379
                        Α1
                                                                  20030716
                                20020801
                                         WO 2002-US1276
     WO 2002058711
                         A1
            AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
             CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
             GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
             LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT,
             RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US,
             UZ, VN, YU, ZA, ZW
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             BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
     WO 2005018544
                         A2
                                20050303
                                          WO 2004-US23099
     WO 2005018544
                         Α3
                                20050609
     WO 2005018544
                         C1
                                20051110
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             NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY,
             TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
         RW: BW, GH, GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM,
             AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK,
             EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE,
             SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE,
             SN, TD, TG
     US 2006020128
                         A1
                               20060126
                                           US 2005-218044
                                                                  20050831
PRIORITY APPLN. INFO.:
                                           US 2001-261911P
                                                              P 20010116
                                           WO 2002-US1276
                                                              A2 20020115
                                           US 2003-621027
                                                               A 20030716
ED
     Entered STN: 18 Jun 2004
     This invention provides a composition comprising an effective amount of glucan
AB
     capable of enhancing efficacy of antibodies. This invention further
     provides the above compns. and a pharmaceutically acceptable carrier.
     This invention also provides a method for treating a subject with cancer
     comprising administrating the above-described composition to the subject.
     invention provides a composition comprising effective amount of glucan capable
of
     enhancing efficacy of vaccines. This invention also provides a method of
     treating a subject comprising administrating the above pharmaceutical
     composition to the subject. This invention provides a composition comprising
     effective amount of glucan capable of enhancing efficacy of natural
     antibodies. This invention provides a composition comprising effective amount
of
     glucan capable of enhancing host immunity. This invention also provides a
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Searched by Barb O'Bryen, STIC 2-2518

composition comprising effective amount of glucan capable of enhancing the

action

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of an agent in preventing tissue rejection.
     ICM A61K031-739
TC
     ICS A61K039-395; A61K031-715
INCL 514054000; 424143100
     15-2 (Immunochemistry)
     Section cross-reference(s): 1, 63
     Antibodies and Immunoglobulins
IT
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (528; antitumor vaccine therapy-enhancing glucan)
     Antibodies and Immunoglobulins
IT
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (C225; antitumor vaccine therapy-enhancing glucan)
     Antibodies and Immunoglobulins
IT
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (R24; antitumor vaccine therapy-enhancing glucan)
     Brain, neoplasm
IT
     Digestive tract, neoplasm
     Gene therapy
     Hodgkin's disease
       Immunostimulants
     Leukemia
     Liver, neoplasm
     Lung, neoplasm
     Mammary gland, neoplasm
     Melanoma
     Molecular weight distribution
     Ovary, neoplasm
     Plasmids
     Stomach, neoplasm
       T cell (lymphocyte)
     Transplant and Transplantation
        (antitumor vaccine therapy-enhancing glucan)
     Antibodies and Immunoglobulins
     RL: PAC (Pharmacological activity); THU (Therapeutic
     use); BIOL (Biological study); USES (Uses)
        (antitumor vaccine therapy-enhancing glucan)
     Antibodies and Immunoglobulins
     RL: PAC (Pharmacological activity); THU (Therapeutic
     use); BIOL (Biological study); USES (Uses)
        (monoclonal; antitumor vaccine therapy-enhancing glucan)
L188 ANSWER 35 OF 63 CAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER:
                         2004:791569 CAPLUS
DOCUMENT NUMBER:
                         142:148164
TITLE:
                         Immune stimulating properties of a novel
                         polysaccharide from the medicinal plant Tinospora
                         cordifolia
                         Raveendran Nair, P. K.; Rodriguez, Sonia;
AUTHOR (S):
                         Ramachandran, Reshma; Alamo, Arturo; Melnick, Steven
                         J.; Escalon, Enrique; Garcia, Pedro I.; Wnuk,
                         Stanislaw F.; Ramachandran, Cheppail
CORPORATE SOURCE:
                         Research Institute, Miami Children's Hospital, Miami,
                         FL, 33155, USA
                         International Immunopharmacology (2004),
SOURCE:
                                                                   4(13),
                         1645-1659
                         CODEN: IINMBA; ISSN: 1567-5769
PUBLISHER:
                         Elsevier B.V.
                         Journal
DOCUMENT TYPE:
LANGUAGE:
                         English
    Entered STN: 29 Sep 2004
```

An  $\alpha$ -D-glucan (RR1) composed of (1 $\rightarrow$ 4) linked back bone and AΒ (1→6) linked branches with a mol. mass of >550 kDa and exhibiting unique immune stimulating properties is isolated and characterized from the medicinal plant Tinospora cordifolia. This novel polysaccharide is noncytotoxic and nonproliferating to normal lymphocytes as well as tumor cell lines at 0-1000 μg/mL. It activated different subsets of the lymphocytes such as natural killer (NK) cells (331%), T cells (102%), and B cells (39%) at 100 μg/mL concentration The significant activation of NK cells is associated with the dose-dependent killing of tumor cells by activated normal lymphocytes in a functional assay. Immune activation by RR1 in normal lymphocytes elicited the synthesis of interleukin  $(IL)-1\beta$  (1080 pg/mL), IL-6 (21,833 pg/mL), IL-12 p70 (50.19 pg/mL), IL-12 p40 (918.23 pg/mL), IL-18 (27.47 pg/mL), IFN- $\gamma$  (90.16 pg/mL), tumor necrosis factor (TNF)- $\alpha$  (2225 pg/mL) and monocyte chemoattractant protein (MCP)-1 (2307 pg/mL) at 100 µg/mL concentration, while it did not induce the production of IL-2, IL-4, IL-10, interferon (IFN)  $-\alpha$  and TNF- $\beta$ . The cytokine profile clearly demonstrates the Th1 pathway of T helper cell differentiation essential for cell mediated immunity, with a self-regulatory mechanism for the control of its overprodn. RR1 also activated the complement alternate pathway, demonstrated by a stepwise increase in C3a des Arg components. Incidentally, RR1 stimulation did not produce any oxidative stress or inducible nitric oxide synthase (iNOS) in the lymphocytes or any significant increase in nitric oxide production The water solubility, high mol. mass, activation of lymphocytes especially NK cells, complement activation, Th1 pathway-associated cytokine profile, together with a low level of nitric oxide synthesis and absence of oxidative stress confer important immunoprotective potential to this novel  $\alpha$ -D-glucan. 1-7 (Pharmacology) CC Section cross-reference(s): 11 T cell (lymphocyte) IT (helper cell/inducer, TH1; immune stimulating properties of a novel polysaccharide from the medicinal plant Tinospora cordifolia) IT B cell (lymphocyte) Cell activation Human Immunostimulants Immunostimulation T cell (lymphocyte) Tinospora cordifolia (immune stimulating properties of a novel polysaccharide from the medicinal plant Tinospora cordifolia) 9074-78-6,  $\alpha$ -D- Glucan TΤ RL: ADV (Adverse effect, including toxicity); NPO (Natural product occurrence); PAC (Pharmacological activity); PRP (Properties); BIOL (Biological study); OCCU (Occurrence) (immune stimulating properties of a novel polysaccharide from the medicinal plant Tinospora cordifolia) REFERENCE COUNT: 62 THERE ARE 62 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT L188 ANSWER 36 OF 63 CAPLUS COPYRIGHT 2006 ACS on STN ACCESSION NUMBER: 2003:796878 CAPLUS DOCUMENT NUMBER: 139:306530 TITLE: Flt3-ligand for enhancing immune response of vaccine against cancer, allergy and infection INVENTOR (S): Mckenna, Hilary J.; Liebowitz, David N.; Maliszewski, Charles R.

Immunex Corporation, USA

PATENT ASSIGNEE(S):

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SOURCE:
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PCT Int. Appl., 96 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

	PA	rent	NO.			KIND DATE				1	APPL	ICAT		DATE						
	• • •				A2 20031009 A3 20040624			1	WO 2	003-1	US97	20030326								
		W:	ΑE,	AG,	AL,	AM,	AT,	AU,	AZ,	BA,	BB,	BG,	BR,	BY,	ΒZ,	CA,	CH,	CN,		
			CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	ES,	FI,	GB,	GD,	GE,	GH,		
			GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KΡ,	KR,	ΚZ,	LC,	LK,	LR,		
			LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NI,	NO,	NZ,	OM,		
			PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	ТJ,	TM,	TN,	TR,	TT,		
			TZ,	UA,	ŪĠ,	US,	UΖ,	VC,	VN,	YU,	ZA,	ZM,	zw							
		RW:	GH,	GM,	ΚE,	LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	ŪĠ,	ZM,	ZW,	AM,	ΑZ,	BY,		
			KG,	ΚŻ,	MD,	RU,	ТJ,	TM,	ΑT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	EE,	ES,		
			FΙ,	FR,	GB,	GR,	HU,	ΙE,	IT,	LU,	MC,	NL,	PT,	RO,	SE,	SI,	SK,	TR,		
			BF,	ВJ,	CF,	CG,	CI,	CM,	GΑ,	GN,	GQ,	GW,	ML,	MR,	NE,	SN,	TD,	TG		
	CA 2480128							2003	1009	CA 2003-2480128						20030326				
	US 2004022760							2004	0205	1	US 2	003-	4013	20030326						
	EP 1487477					A2	20041222			EP 2003-721501						20030326				
		R:	ΑT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,		
			ΙE,	SI,	LT,	LV,	FI,	RO,	MK,	CY,	AL,	TR,	BG,	CZ,	EE,	HU,	SK			
JP 2005528373							T2 20050922				JP 2	003-	5805	20030326						
PRIORITY APPLN. INFO.:										US 2002-368263P					1	P 20020326				
										1	US 2002-427835P					P 20021119				
					1	WO 2	003-1	US97'	73	I	W 2	0030	326 /							

ED Entered STN: 10 Oct 2003

The present invention relates to methods of using Flt3-ligand (Flt3-L) in immunization protocols to enhance immune responses against vaccine antigens. Embodiments include administering Flt3-ligand prior to immunizing a subject with a vaccine, wherein the vaccine comprises at least one antigen formulated in one or more adjuvants. Methods of treating and preventing cancer, allergy and infection using Flt3-ligand immunization protocols are also provided. Methods of using Flt3-ligand immunization protocols for in vivo evaluation of antigens and adjuvants are also provided.

IC ICM C12N

CC 15-2 (Immunochemistry)

Section cross-reference(s): 63

IT AIDS (disease)

Actinomyces israelii African swine fever virus

Allergy

Antitumor agents

Arenaviridae

Aspergillus fumigatus

Astrovirus

Bacteroides

Birnaviridae

Blastomyces dermatitidis

Borrelia burgdorferi

Bunyaviridae

Bunyavirus

CD4-positive T cell

CD8-positive T cell

Calicivirus

Candida albicans Chlamydia trachomatis Clostridium perfringens Clostridium tetani Coccidioides immitis Coronaviridae Coronavirus Corynebacterium Corynebacterium diphtheriae Cryptococcus neoformans Cytomegalovirus Dengue virus Ebola virus Enterobacter aerogenes Enterococcus faecalis Enterovirus **Epitopes** Equine encephalosis virus Erysipelothrix rhusiopathiae Eubacteria Filoviridae Flaviviridae Fusobacterium nucleatum Granulicatella adiacens Hantaan virus Hantavirus Helicobacter pylori Hepadnaviridae Hepatitis A virus Hepatitis B virus Herpesviridae Histoplasma capsulatum Human Human adenovirus Human coxsackievirus Human echovirus Human herpesvirus 1 Human herpesvirus 2 Human herpesvirus 3 Human immunodeficiency virus 3 Human parainfluenza virus Human poliovirus Immunization

# Immunostimulants

Immunotherapy

Infection
Influenza virus
Iridoviridae
Klebsiella pneumoniae
Legionella pneumophila
Leishmania
Leptospira
Listeria monocytogenes
Measles virus
Melanoma
Microparticles
Microspheres
Mumps virus
Mus
Mycobacterium

Mycobacterium avium Mycobacterium gordonae Mycobacterium intracellulare Mycobacterium kansasii Mycobacterium tuberculosis Mycosis Nairovirus Nanoparticles Neisseria gonorrhoeae Neisseria meningitidis Norwalk virus Orbivirus Orthomyxoviridae Papillomavirus Papovaviridae Paramyxoviridae Parvoviridae Parvovirus Pasteurella multocida Pathogen Phlebovirus Picornaviridae Plasmodium falciparum Plasmodium gonderi Plasmodium malariae Plasmodium vivax Polyomavirus Poxviridae Protein sequences Rabies virus Reoviridae Respiratory syncytial virus Retroviridae Rhabdoviridae Rhinovirus Rotavirus Rubella virus Sarcosporida Schistosoma Staphylococcus aureus Streptobacillus moniliformis Streptococcus agalactiae Streptococcus anaerobius Streptococcus bovis Streptococcus group A Streptococcus group B Streptococcus pneumoniae Streptococcus pyogenes Taenia saginata Taenia solium Toqaviridae Treponema pallidum Treponema pallidum pertenue Trichinella Trichomonas Trypanosoma Vaccines Vaccinia virus Variola virus Vesicular stomatitis virus

10/630143 Page 68 Jones

Yellow fever virus

(Flt3-ligand for enhancing immune response of vaccine against cancer, allergy and infection)

IT Immunostimulants

> (adjuvants, DRVs; Flt3-ligand for enhancing immune response of vaccine against cancer, allergy and infection)

IT Immunostimulants

> (adjuvants, Freund's incomplete; Flt3-ligand for enhancing immune response of vaccine against cancer, allergy and infection)

IT Immunostimulants

> (adjuvants, Freund's; Flt3-ligand for enhancing immune response of vaccine against cancer, allergy and infection)

Immunostimulants IT

(adjuvants, ISCOMs; Flt3-ligand for enhancing immune response of vaccine against cancer, allergy and infection)

Immunostimulants TT

(adjuvants; Flt3-ligand for enhancing immune response of vaccine against cancer, allergy and infection)

TT T cell (lymphocyte)

(cytotoxic; Flt3-ligand for enhancing immune response of vaccine against cancer, allergy and infection)

ΙT T cell (lymphocyte)

(helper cell; Flt3-ligand for enhancing immune response of vaccine against cancer, allergy and infection)

9012-72-0, Glucan TT

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (algal; Flt3-ligand for enhancing immune response of vaccine against cancer, allergy and infection)

L188 ANSWER 37 OF 63 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER:

2002:465829 CAPLUS

DOCUMENT NUMBER:

137:37673

TITLE:

Immune system stimulation with agents stimulating T

lymphocytes

INVENTOR(S):

Graus, Yvo Maria Franciscus; Smit, Hobbe Friso; Osterhaus, Albertus Dominicus Marcellinus Erasmus;

Hageman, Robert Johan Joseph

PATENT ASSIGNEE(S):

Nutricia N.V., Neth. PCT Int. Appl., 25 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

SOURCE:

English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT	NO.			KIND		DATE			APPLICATION NO.						DATE			
WO 2002	0477	03		A2 20020620			1	WO 2	001-1		20011210							
WO 2002	A3 20020912																	
WO 2002	C1		2004	0513														
W:	ΑE,	AG,	AL,	AM,	AT,	AU,	ΑZ,	BA,	BB,	BG,	BR,	BY,	ΒZ,	CA,	CH,	CN,		
	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	ES,	FI,	GB,	GD,	GE,	GH,		
	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KP,	KR,	ΚZ,	LC,	LK,	LR,		
	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	ΜZ,	NO,	NZ,	OM,	PH,		
	PL,	PT,	RO,	RU,	SD,	SE,	SG,	SI,	SK,	SL,	ТJ,	TM,	TN,	TR,	TT,	TZ,		
	UA,	ŪĠ,	US,	UΖ,	VN,	ΥU,	ZA,	ZM,	zw									
RW:	GH,	GM,	KE,	LS,	MW,	MZ,	SD,	SL,	SZ,	ΤŹ,	UG,	ZM,	ZW,	AM,	ΑZ,	BY,		
	KG,	KZ,	MD,	RU,	TJ,	TM,	ΑT,	BE,	CH,	CY,	DE,	DK,	ES,	FI,	FR,	GB,		
	GR,	IE,	IT,	LU,	MC,	NL,	PT,	SE,	TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GA,		
	GN,	GQ,	GW,	ML,	MR,	NE,	SN,	TD,	TG									

20020815 US 2000-734389 20001211 US 2002110606 A1 20031014 US 6632459 B2 AU 2002025512 Α5 20020624 AU 2002-25512 20011210 20040520 20030702 US 2004097584 **A1** US 2003-612242 20001211 PRIORITY APPLN. INFO.: US 2000-734389 Α 20011210 WO 2001-NL896 W

OTHER SOURCE(S): MARPAT 137:37673

ED Entered STN: 21 Jun 2002

- The present invention relates to a preparation for stimulating or enhancing an immune system comprising one or more agents that stimulate T-lymphocytes vivo. Such a preparation can be used in the prophylaxis and/or treatment of a medical condition. The invention further relates to a preparation for use in a pharmaceutical or food product and to a preparation for medical use. Example stimulants include ascorbic acid, N-acetylcysteine, chlorogenic acid and derivs., and plant exts.
- IC ICM A61K035-78
  - ICS A61P037-04
- CC 63-6 (Pharmaceuticals)
  - Section cross-reference(s): 1, 15, 17
- IT Immunostimulants

(adjuvants; immune system stimulation with agents stimulating T lymphocytes)

IT Acanthopanax senticosus Achyrocline satureioides Aconitum officinalis

Aconitum officinalis
Angelica acutiloba

Anti-infective agents

Antibacterial agents

Antioxidants

Antitumor agents

Antiviral agents

Aristolochia officinalis

Arnica montana

Asteraceae

Astragalus gummifer

Astragalus membranaceus

Astragalus penduliflorus

Avena sativa

Bambusa vulgaris

Baptisia tinctoria

Betula

Beverages

Bryonia dioica

Butia capitata

Calendula officinalis

Carthamus tinctorius

Crataegus

Cynanchum vincetoxicum

Cynara scolymus

Echinacea

Echinacea angustifolia

Echinacea pallida

Echinacea purpurea

Eupatorium cannabinum

Flammulina velutipes

Food

## Immunostimulants

Larix occidentalis

Matricaria recutita

Panax pseudoginseng

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Parasiticides
     Paullinia cupana
     Petasites
     Phoenix
     Plantago
     Sambucus
     Silene vulgaris
       T cell (lymphocyte)
     Taraxacum officinale
     Thuja occidentalis
     Triticum aestivum
     Vaccines
     Viscum album
        (immune system stimulation with agents stimulating T
        lymphocytes)
     7440-66-6, Zinc, biological studies
                                            9041-22-9, \beta- Glucan
IT
     RL: MOA (Modifier or additive use); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (immune system stimulation with agents stimulating T
        lymphocytes)
L188 ANSWER 38 OF 63 CAPLUS COPYRIGHT 2006 ACS on STN
                         2001:248195 CAPLUS
ACCESSION NUMBER:
DOCUMENT NUMBER:
                         135:70891
                         Immunostimulant oxidized \beta-glucan conjugates
TITLE:
                         Cross, G. G.; Jennings, H. J.; Whitfield, D. M.;
AUTHOR (S):
                         Penney, C. L.; Zacharie, B.; Gagnon, L.
                         National Research Council, Ottawa, ON, K1A OR6, Can.
CORPORATE SOURCE:
                         International Immunopharmacology (2001), 1(3), 539-550
SOURCE:
                         CODEN: IINMBA; ISSN: 1567-5769
PUBLISHER:
                         Elsevier Science B.V.
                         Journal
DOCUMENT TYPE:
LANGUAGE:
                         English
     Entered STN: 08 Apr 2001
ED
     β-Glucans are polysaccharides that act as nonspecific immune system
AB
     stimulants. However, many β-Glucans are sparingly soluble in water.
     This work describes an oxidative procedure, which solubilizes the
     β-glucan from Saccharomyces cerevisiae and maintains its
     immunostimulatory properties. Furthermore, the carboxylates at the site
     of oxidation allow for the conjugation of small mol. immunostimulants. Both
     the parent oxidized \beta-glucan and its conjugates with
     O-β-alanyl-5-[6-(N,N'-dimethylamino)purin-9-yl]pentanol stimulate
     cytotoxic T-lymphocytes (CTLs), B cells and macrophages. In addition, they
     both stimulate natural killer (NK) cells, a property which the small mol.
     purine does not possess.
     1-7 (Pharmacology)
CC
     T cell (lymphocyte)
TT
        (cytotoxic, stimulation; immunostimulant oxidized β-glucan
        conjugates)
IT
     Immunostimulants
     Saccharomyces cerevisiae
        (immunostimulant oxidized β-glucan conjugates)
     Polysaccharides, biological studies
TТ
     RL: BAC (Biological activity or effector, except adverse); BSU
     (Biological study, unclassified); BIOL (Biological study)
        (immunostimulant oxidized \beta- glucan conjugates)
TΤ
     9041-22-9D, β- Glucan, conjugates 194225-61-1D, conjugates
     with β- glucans
     RL: BAC (Biological activity or effector, except adverse); BSU
     (Biological study, unclassified); BIOL (Biological study)
```

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(immunostimulant oxidized \beta- glucan conjugates)
                             THERE ARE 41 CITED REFERENCES AVAILABLE FOR THIS
                       41
REFERENCE COUNT:
                             RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
L188 ANSWER 39 OF 63 CAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER: 1999:350607 CAPLUS
DOCUMENT NUMBER:
                       131:14825
                       A method of increasing nucleic acid synthesis with
TITLE:
                       ultrasound
                       Unger, Evan C.; McCreery, Thomas; Sadewasser, David
INVENTOR(S):
                       ImaRx Pharmaceutical Corp., USA
PATENT ASSIGNEE(S):
                       PCT Int. Appl., 124 pp.
SOURCE:
                       CODEN: PIXXD2
DOCUMENT TYPE:
                       Patent
                       English
LANGUAGE:
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
                       KIND DATE
                                        APPLICATION NO.
                                                              DATE
    PATENT NO.
                                         -----
                                                                _____
     ______
                        ----
                              19990527 WO 1998-US23843 19981111
     WO 9925385
                        A1
        W: AU, CA, JP
        RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
            PT, SE
                              19990607
                                          AU 1999-13906
                                                                19981111
    AU 9913906
                        A1
                                          US 1997-971540
                                                           A 19971117
PRIORITY APPLN. INFO.:
                                                           W 19981111
                                          WO 1998-US23843
OTHER SOURCE(S):
                       MARPAT 131:14825
    Entered STN: 08 Jun 1999
     The present invention is directed to a method of increasing nucleic acid
AB
     synthesis in a cell comprising administering to the cell a therapeutically
     effective amount of ultrasound for a therapeutically effective time such
     that said administration of said ultrasound results in said in reased
     nucleic acid synthesis. The nucleic acid sequence may comprise an
     endogenous sequence or an exogenous sequence. In particular, the
     invention is directed to increasing the expression of stress proteins and
     repair proteins.
    ICM A61K048-00
IC
     ICS A61H001-00
     3-1 (Biochemical Genetics)
CC
     Section cross-reference(s): 1, 6, 9, 11, 13, 14
IT
     T cell (lymphocyte)
        (killer cell; method of increasing nucleic acid synthesis with
       ultrasound)
     50-69-1D, Ribose, polymers containing 50-99-7D, Glucose, polymers
IT
     containing 57-09-0, CTAB 57-10-3, Palmitic acid, biological studies
     57-11-4, Octadecanoic acid, biological studies 57-48-7D, Fructose,
     polymers containing 57-88-5, Cholesterol, biological studies
     Cholesterol, derivs. 57-88-5D, Cholesterol, ester and salt 58-73-1,
     DPH 58-86-6D, Xylose, polymers containing 59-23-4D, Galactose, polymers
     containing 65-42-9D, Lyxose, polymers containing 87-79-6D, Sorbose,
polymers
     containing 112-80-1, 9-Octadecenoic acid (9Z)-, biological studies
     114-04-5D, Neuraminic acid, polymers containing 124-30-1, Stearylamine
     147-81-9D, Arabinose, polymers containing 506-32-1, Arachidonic acid
     526-95-4D, Gluconic acid, polymers containing 685-73-4D, Galacturonic acid,
     polymers containing 926-63-6 1122-58-3, DMAP 1256-86-6, Cholesterol
     sulfate 1398-61-4, Chitin 1398-61-4D, Chitin, derivative
     1510-21-0, Cholesterol hemisuccinate 1758-51-6D, Erythrose, polymers
     containing 2152-76-3D, Idose, polymers containing 2390-68-3, DDAB
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2462-63-7,

```
2644-64-6, Dipalmitoylphosphatidylcholine
                                                        3416-24-8D,
    Glucosamine, polymers containing 3458-28-4D, Mannose, polymers containing
     3700-67-2, Dimethyldioctadecylammonium bromide
                                                    4235-95-4, DOPC
               4458-31-5
                            4539-70-2, Distearoylphosphatidylcholine
     4345-03-3
     5556-48-9D, Ribulose, polymers containing 5962-29-8D, Xylulose, polymers
    containing 5987-68-8D, Altrose, polymers containing 6038-51-3D, Allose,
    polymers containing 6556-12-3D, Glucuronic acid, polymers containing
     6561-76-8, DCPE 6814-36-4D, Mannuronic acid, polymers containing
     7439-95-4, Magnesium, biological studies 7440-66-6, Zinc, biological
             7440-70-2, Calcium, biological studies 7535-00-4D,
    Galactosamine, polymers containing 9000-07-1, Carrageenan 9000-69-5,
     Pectin 9002-88-4D, Polyethylene, derivs. 9002-89-5D, Polyvinyl
     alcohol, derivs. 9003-07-0D, Polypropylene, derivs. 9003-39-8,
     Polyvinylpyrrolidone 9003-39-8D, Polyvinylpyrrolidone, derivative
               9004-34-6, Cellulose, biological studies 9004-54-0, Dextran,
    biological studies 9004-61-9, Hyaluronic acid 9004-61-9D, Hyaluronic
     acid, derivative 9004-65-3, Hydroxypropyl methylcellulose 9005-32-7,
    Alginic acid 9005-79-2, Glycogen, biological studies
                                                            9005-82-7,
             9007-27-6, Chondroitin 9012-36-6, Agarose 9012-72-0D,
    Amvlose
     Glucan, derivs. 9013-95-0, Levan 9014-63-5D, Xylan, derivs.
     9036-88-8D, Mannan, derivs. 9037-22-3, Amylopectin 9037-55-2D,
    Galactan, derivs. 9037-90-5D, Fructan, derivs. 9046-38-2D,
    Galacturonan, derivs. 9046-40-6, Pectic acid 9057-02-7, Pullulan
     9060-75-7D, Arabinan, derivs. 9072-19-9, Fucoidan 15769-56-9D,
    Guluronic acid, polymers containing 17598-81-1D, Tagatose, polymers
containing
     18656-38-7, Dimyristoylphosphatidylcholine
                                                18656-40-1,
     Dilauroylphosphatidylcholine 19163-87-2D, Gulose, polymers containing
     19600-01-2, Ganglioside GM2 19698-29-4, Dipalmitoylphosphatidic acid
                 20255-95-2, DMPE 23140-52-5D, Psicose, polymers containing
     20064-29-3
                 24529-88-2 25322-68-3D, Polyethylene glycol, alcs.
     24305-42-8
     25322-68-3D, Polyethylene glycol, derivative 25322-68-3D, derivs.
     25525-21-7D, Glucaric acid, polymers containing 29884-64-8D, Threose,
    polymers containing 30077-17-9D, Talose, polymers containing 37331-28-5, Pustulan 37758-47-7, Ganglioside GM1 40031-31-0D, Erythrulose,
    polymers containing 60495-58-1, Galactocarolose 64612-25-5D, Fucan, derivs. 67896-63-3, Dipentadecanoylphosphatidylcholine 68354-92-7
                 68737-67-7, Dioleoylphosphatidylcholine 69992-87-6, Keratan
     68354-99-4
                 75634-40-1, Dermatan 76822-97-4 78543-25-6 83554-62-5
     73294-85-6
     106392-12-5, Pluronic 106392-12-5D, Pluronic, acid and alc. derivs.
                  115534-33-3, TMADPH 124050-77-7, Transfectam
     108032-13-9
                                                                  124076-29-5
     127512-30-5 128835-92-7, Lipofectin 137056-72-5, DC-Chol
     144189-73-1, DOTAP
                        145035-97-8, Dipalmitoylphosphatidylethanolamine-PEG
     145310-87-8, Transfectace 153312-64-2, DMRIE 158571-62-1,
                   161293-59-0 161441-83-4 165467-64-1, DOHME
    Lipofectamine
     168479-03-6, DOSPA
                         182919-20-6 183283-19-4, EDMPC 186198-32-3
                         201491-17-0, Cytofectin 214206-92-5
     199171-54-5, DLRIE
                                                                  214206-94-7
                  225940-36-3
                                225940-37-4
                                             225940-38-5
                                                           225940-42-1
    225940-35-2
    225940-43-2
    RL: BPR (Biological process); BSU (Biological study, unclassified); BUU
     (Biological use, unclassified); THU (Therapeutic use); BIOL (Biological
     study); PROC (Process); USES (Uses)
        (carrier; method of increasing nucleic acid synthesis with ultrasound)
                               THERE ARE 12 CITED REFERENCES AVAILABLE FOR THIS
REFERENCE COUNT:
                         12
                               RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
```

L188 ANSWER 40 OF 63 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 1994:645530 CAPLUS

DOCUMENT NUMBER: 121:245530

Immunorestorative effect of glucan immunomodulator on TITLE:

guinea pigs with experimental ascariosis AUTHOR (S): Soltys, J.; Benkova, M.; Boroskova, Z. Parasitological Inst., SAS, Kosice, 040 01, Slovakia CORPORATE SOURCE: Veterinary Immunology and Immunopathology (1994), SOURCE: 42 (3-4), 379-88 CODEN: VIIMDS; ISSN: 0165-2427 DOCUMENT TYPE: Journal LANGUAGE: English Entered STN: 26 Nov 1994 The immunorestorative effect of glucan immunomodulator, combined with porcine Ig and zinc (GI) on T- and B-lymphocytes and peritoneal macrophage phagocytic ability was studied in guinea pigs with exptl. ascariosis (Ascaris suum) after a cyclophosphamide (CY)-evoked immunosuppression. During the migration phase of A. suum infection GI exerted a significant restorative effect on the CY-reduced percentage occurrence of T- and B-cell populations in the mesenteric, mediastinal and hepatic lymph nodes and spleen of A. suum hosts. On the contrary, it did not influence the CY-suppressed phagocytic activity and index of phagocytic activity of the peritoneal macrophages. The protective effect of the GI evaluated by the reduction in the number of migrating ascarid larvae in the lungs of guinea pigs after immunosuppression with CY and administration of GIZ was 14.46% higher, compared with the suppressed and infected group without administration of GI. 1-7 (Pharmacology) CC IT Immunoglobulins RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (combination with zinc and glucan immunomodulator; immunorestorative effect of glucan immunomodulator on guinea pigs with exptl. ascariosis) IT Immunostimulants (immunorestorative effect of glucan immunomodulator on guinea pigs with exptl. ascariosis) Lymphocyte TT (T-cell, immunorestorative effect of glucan immunomodulator on guinea pigs with exptl. ascariosis) 7440-66-6, Zinc, biological studies TT RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (combination with porcine Ig and glucan immunomodulator; immunorestorative effect of glucan immunomodulator on guinea pigs with exptl. ascariosis) IT 9012-72-0, D-Glucan RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (immunorestorative effect of glucan immunomodulator on guinea pigs with exptl. ascariosis)

L188 ANSWER 41 OF 63 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN DUPLICATE 1

ACCESSION NUMBER: 2003:348988 BIOSIS DOCUMENT NUMBER: PREV200300348988

TITLE: Immunopotentiation of intraepithelial lymphocytes

in the intestine by oral administrations of beta-

glucan.

AUTHOR (S): Tsukada, Chika; Yokoyama, Hisashi; Miyaji, Chikako;

Ishimoto, Yuiko; Kawamura, Hiroki; Abo, Toru (Reprint

Department of Immunology, Niigata University School of CORPORATE SOURCE:

Medicine, Niigata, 951-8510, Japan

immunol2@med.niigata-u.ac.jp

Cellular Immunology, (January 2003) Vol. 221, No. 1, pp. SOURCE:

1-5. print.

CODEN: CLIMB8. ISSN: 0008-8749.

Article DOCUMENT TYPE: English LANGUAGE:

ENTRY DATE: Entered STN: 30 Jul 2003

Last Updated on STN: 30 Jul 2003

ABSTRACT: Mice were orally administered with beta-glucan,

isolated from baker's yeast, daily for one week (25 mg/day/mouse) and several immunoparameters in the digestive tract were examined. The most prominent change was an increase in the number of intraepithelial lymphocytes (IEL) in the intestine, although the number of lymphocytes in the liver remained unchanged. The absolute number of both alphabetaT cells and gammadeltaT cells expressing CD8 antigens increased among IEL in the intestine. Primarily, liver lymphocytes showed a spontaneous production of Type 0 cytokine (simultaneous production of IFNgamma and IL-4) while IEL did not produce any cytokines without stimulation. However, mice administered with beta-

\*\*\*glucan\*\*\* produced Type 1 cytokine, namely, production of IFNgamma alone.

These results suggest that beta-glucan may be an important potentiator for mucosal immunity in the digestive tract.

CONCEPT CODE: Cytology - General 02502 Cytology - Plant 02504

Cytology - Animal Biochemistry studies - Proteins, peptides and amino acids

10064

Pathology - Therapy 12512

Digestive system - Physiology and biochemistry 14004

Blood - Blood and lymph studies 15002

02506

Blood - Blood cell studies 15004

Endocrine - General 17002 Pharmacology - General 22002

Pharmacology - Immunological processes and allergy 22018

Immunology - General and methods 34502

Major Concepts INDEX TERMS:

Cell Biology; Digestive System (Ingestion and

Assimilation); Immune System (Chemical Coordination and

Homeostasis); Pharmacology

INDEX TERMS: Parts, Structures, & Systems of Organisms

alpha-beta T cell: blood and

lymphatics, immune system; digestive tract: digestive

system; gamma-delta T cell: blood

and lymphatics, immune system; intestine: digestive

system; intraepithelial lymphocyte: blood and

lymphatics, immune system; liver: digestive system

Chemicals & Biochemicals INDEX TERMS:

CD8: antigen; IFN-gamma [interferon-gamma]: cytokine;

IL-4 [interleukin-4]: cytokine; beta-

glucan: immunologic-drug, oral administration

Miscellaneous Descriptors INDEX TERMS:

immunopotentiation; mucosal immunity

ORGANISM: Classifier

> Ascomycetes 15100

Super Taxa

Fungi; Plantae Organism Name

baker's yeast (common)

Taxa Notes

Fungi, Microorganisms, Nonvascular Plants, Plants

ORGANISM: Classifier

Muridae 86375

Super Taxa

Rodentia; Mammalia; Vertebrata; Chordata; Animalia

Organism Name

mouse (common): strain-C57BL/6

Taxa Notes

Animals, Chordates, Mammals, Nonhuman Vertebrates,

Nonhuman Mammals, Rodents, Vertebrates

REGISTRY NUMBER: 9041-22-9 (beta-glucan)

L188 ANSWER 42 OF 63 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on

STN DUPLICATE 4

ACCESSION NUMBER: 1999:510610 BIOSIS DOCUMENT NUMBER: PREV199900510610

TITLE: Enhancement of cytotoxic T lymphocyte

activity by gel-forming (1fwdarw3)-beta-D-

glucan, SSG.

AUTHOR(S): Adachi, Y.; Ishikawa, M.; Ohno, N.; Yadomae, T. [Reprint

authorl

CORPORATE SOURCE: Laboratory of Immunopharmacology of Microbial Products,

Tokyo University of Pharmacy and Life Science, 1432-1

Horinouchi, Hachioji, Tokyo, 192-0392, Japan

SOURCE: Pharmaceutical and Pharmacological Letters, (July 1999)

Vol. 9, No. 1, pp. 14-17. print.

ISSN: 0939-9488.

DOCUMENT TYPE: Article LANGUAGE: English

ENTRY DATE: Entered STN: 3 Dec 1999

Last Updated on STN: 3 Dec 1999

ABSTRACT: The effect of the (1 fwdarw 3)-beta-D-glucan, SSG, on generation of alloreactive cytotoxic T lymphocytes (CTL)

in a primary murine mixed lymphocyte reaction (MLR) was examined. Cytotoxic activity induced by SSG is dependent on concentration of SSG and is restricted to allogenic antigen. To determine molecular mechanism of cytotoxicity induced by SSG, expression of cell-surface molecules and production of cytokines were examined. SSG augmented expression of IL-12 and IL-15, which might enhance CTL activity and strengthen cellular immunity. Since induction of IFN-gamma in supernatant was dependent on concentration of SSG in MLR, it was suggested that IFN-gamma (nay stimulate macrophages to enhance expression of adhesion molecules. These results suggested that induction of antigen specific CTL in the presence of SSG was mediated by production of IL-12 and IL-15 or expressions of ICAM-1, B7-1 and B7-2 via augmentation of

INF-gamma production.

CONCEPT CODE: Immunology - General and methods 34502 Biochemistry studies - General 10060

> Biophysics - General 10502 Endocrine - General 17002 Pharmacology - General 22002

INDEX TERMS: Major Concepts

Immune System (Chemical Coordination and Homeostasis);

Pharmacology

INDEX TERMS: Parts, Structures, & Systems of Organisms

cytotoxic T lymphocytes: blood and

Searched by Barb O'Bryen, STIC 2-2518



Jones 10/630143 Page 76

lymphatics, immune system Chemicals & Biochemicals INDEX TERMS: (1-3)-beta-D-glucan [SSG]; allogenic antigen; **B7-1**; **B7-2**; ICAM-1 [intercellular adhesion molecule-1]; IFN-gamma [interferon-gamma]; IL-12 [interleukin-12]; IL-15 [interleukin-15] INDEX TERMS: Miscellaneous Descriptors cellular immunity; mixed lymphocyte reaction Classifier ORGANISM: Muridae 86375 Super Taxa Rodentia; Mammalia; Vertebrata; Chordata; Animalia Organism Name murine Taxa Notes Animals, Chordates, Mammals, Nonhuman Vertebrates, Nonhuman Mammals, Rodents, Vertebrates REGISTRY NUMBER: 9051-97-2 ((1-3)-beta-D-glucan 9051-97-2 (SSG) L188 ANSWER 43 OF 63 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on DUPLICATE 5 STN 1996:126183 BIOSIS ACCESSION NUMBER: DOCUMENT NUMBER: PREV199698698318 Pulmonary metastases neutralization and tumor rejection by TITLE: in vivo administration of beta glucan and bispecific antibody. AUTHOR (S): Penna, Christophe; Dean, Phillip A.; Nelson, Heidi [Reprint author] Dep. Surg., Mayo Clinic, 200 First St. S.W., Rochester, MN CORPORATE SOURCE: 55905, USA International Journal of Cancer, (1996) Vol. 65, No. 3, pp. SOURCE: 377-382. CODEN: IJCNAW. ISSN: 0020-7136. DOCUMENT TYPE: Article LANGUAGE: English Entered STN: 27 Mar 1996 ENTRY DATE: Last Updated on STN: 27 Mar 1996 ABSTRACT: Bispecific antibody (BsAb) with specificity for tumor cell surface antigen and the CD3 molecule on T cells can redirect activated T cells to lyse tumor cells. Since the ex vivo expansion and activation of T cells is impractical and ineffective for treating established tumors, we tested whether the \*\*\*immune\*\*\* stimulant beta glucan could in situ-activate T cells, which could secondarily be retargeted with BsAbs to lyse tumor cells. To test for tumor neutralization, C3H/HeN mice were injected i.v. with CI-62 melanoma cells and immediately treated with i.p. beta glucan and/or anti-CD3 (500A2) times anti-p97 (96.5) F(ab')-2 BsAb i.v. Pulmonary metastases were counted 14 days later. To test for tumor rejection and survival in a solid tumor model, mice were injected s.c. and i.p. with CI-62 cells and 7 days later administered \*\*\*beta\*\*\* glucan i.p. and/or F(ab')-2 BsAb i.v. In the neutralization model, there was a significant reduction in the number of metastases in the beta glucan + BsAb group, as compared with controls, and with beta glucan alone. In the

incidence of s.c. tumors as compared with control, with BsAb alone and with glucan alone. It also prolonged survival of

established tumor model, beta glucan + BsAb reduced the

\*\*\*beta\*\*\*

tumor-bearing mice compared with control, BsAb alone and beta alone. We conclude that T cells can be \*\*\*glucan\*\*\* activated in vivo by glucan and retargeted with F(ab')-2 BsAb. Cytology - Animal 02506 CONCEPT CODE: Biochemistry studies - Proteins, peptides and amino acids 10064 Biochemistry studies - Carbohydrates 10068 Pathology - Therapy 12512 Metabolism - Carbohydrates 13004 Metabolism - Proteins, peptides and amino acids 13012 Blood - Blood cell studies 15004 Blood - Lymphatic tissue and reticuloendothelial system 15008 Respiratory system - Pathology 16006 Integumentary system - Pathology 18506 Pharmacology - Clinical pharmacology Neoplasms - Immunology 24003 Neoplasms - Pathology, clinical aspects and systemic effects 24004 Neoplasms - Biochemistry 24006 Neoplasms - Therapeutic agents and therapy Immunology - Immunopathology, tissue immunology INDEX TERMS: Major Concepts Blood and Lymphatics (Transport and Circulation); Cell Biology; Immune System (Chemical Coordination and Homeostasis); Integumentary System (Chemical Coordination and Homeostasis); Metabolism; Respiratory System (Respiration); Tumor Biology INDEX TERMS: Chemicals & Biochemicals BETA GLUCAN Miscellaneous Descriptors INDEX TERMS: ACTIVATED T CELL; MELANOMA CELL; POTENTIAL THERAPY; SURVIVAL; TARGETING; TUMOR CELL LYSIS ORGANISM: Classifier 86375 Muridae Super Taxa Rodentia; Mammalia; Vertebrata; Chordata; Animalia Organism Name mouse Taxa Notes Animals, Chordates, Mammals, Nonhuman Vertebrates, Nonhuman Mammals, Rodents, Vertebrates REGISTRY NUMBER: 9041-22-9 (BETA GLUCAN) L188 ANSWER 44 OF 63 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN 2005:119500 BIOSIS ACCESSION NUMBER: PREV200500117957 DOCUMENT NUMBER: Short- and long-term effects of a dietary yeast TITLE: beta-glucan (Macrogard) and alginic acid (Ergosan) preparation on immune response in sea bass (Dicentrarchus labrax). Bagni, M.; Romano, N.; Finoia, M. G.; Abelli, L.; AUTHOR (S): Scapighati, G.; Tiscar, P. G.; Sarti, M.; Marino, G. [Reprint Author] Dept AquacultureInst Res Appl Sea, ICRAM, Via Casalotti CORPORATE SOURCE: 300, I-00166, Rome, Italy g.marino@icram.org

Searched by Barb O'Bryen, STIC 2-2518

pp. 311-325. print.

SOURCE:

Fish & Shellfish Immunology, (April 2005) Vol. 18, No. 4,

Jones 10/630143 Page 78

ISSN: 1050-4648.

DOCUMENT TYPE:

Article

LANGUAGE: ENTRY DATE: English

Entered STN: 23 Mar 2005 Last Updated on STN: 23 Mar 2005

ABSTRACT: The present study investigated the **immunomodulatory** activity of Ergosan, an algal extract containing alginic acid, and Macrogard, a yeast extract containing **beta-glucans**, on innate and specific

immunity in sea bass (Dicentrarchus labrax). Four cycles of experimental feeding using normal fish feed formulation (control group) supplemented with Ergosan (0.5%) or Macrogard (0.1%) were performed at 60-day intervals (15 days of treatment + 45 days of suspension). Serum complement, lysozyme, total proteins and heat shock protein (HSP) concentrations were measured at 15, 30 and 45 days from the end of the first 15-day feeding cycle (short term) and 45 days after the end of each feeding cycle over a 35-week period (long term). The percentage of B- and T-lymphocytes in peripheral blood

leucocytes and gut were measured over long-term trial. Significant elevation (P < 0.05) in serum complement activity occurred in sea bass fed with alginic

acid and glucans, at 15 days from the end of first cycle of treatment. Significant elevation (P < 0.05) in serum lysozyme, gill and liver HSP concentration were observed in the same experimental groups at 30 days from the end of treatment, whereas a significant increase (P < 0.05) of complement activity was only observed in fish that received an Ergosan diet. At 45 days from the end of treatment, complement, lysozyme and HSP concentration did not differ among groups. Over the long-term period, no significant differences were observed in innate and specific immune parameters, survival, growth

performances and conversion index in treated and control fish. A dramatic decrease of both innate and acquired immune parameters was observed during the winter season in all groups, followed by a partial recovery when water temperature increased. Reduction in complement and lysozyme activities was significatively correlated (P < 0.01) to water temperature variation. The results suggested the potential of alginic acid and beta-

\*\*\*glucans\*\*\* to activate some innate immune responses in sea bass, and particularly under conditions of immunodepression related to environmental stress. Copyright 2004 Elsevier Ltd. All rights reserved.

CONCEPT CODE:

Cytology - Animal 02506

Ecology: environmental biology - Animal 07508
Ecology: environmental biology - Oceanography 07512
Biochemistry studies - Proteins, peptides and amino acids
10064

Biochemistry studies - Carbohydrates 10068

Enzymes - General and comparative studies: coenzymes

10802

Food technology - General and methods 13502

Digestive system - Physiology and biochemistry 14004

Blood - Blood and lymph studies 15002

Blood - Blood cell studies 15004

Respiratory system - Physiology and biochemistry 16004 Pharmacology - Immunological processes and allergy 22018

Immunology - General and methods 34502

INDEX TERMS: Major Concepts

Digestive System (Ingestion and Assimilation); Foods; Immune System (Chemical Coordination and Homeostasis);

Marine Ecology (Ecology, Environmental Sciences)

INDEX TERMS: Parts, Structures, & Systems of Organisms

B lymphocyte: blood and lymphatics, immune system;

T lymphocyte: blood and lymphatics,

immune system; gill: respiratory system; gut: digestive system; leukocyte: blood and lymphatics, immune system;

liver: digestive system

Jones 10/630143 Page 79

INDEX TERMS:

Chemicals & Biochemicals

alginic acid [Ergosan]; beta-glucan
[Macrogard]; heat shock protein;

immunostimulants; lysozyme [EC 3.2.1.17]

INDEX TERMS:

Methods & Equipment

experimental feeding cycle: applied and field techniques

INDEX TERMS: Miscellaneous Descriptors

immune response;

immunomodulatory activity; water temperature;

winter

ORGANISM:

Classifier

Osteichthyes 85206

Super Taxa

Pisces; Vertebrata; Chordata; Animalia

Organism Name

Dicentrarchus labrax (species) [sea bass (common)]

Taxa Notes

Animals, Chordates, Fish, Nonhuman Vertebrates,

Vertebrates

REGISTRY NUMBER:

9005-32-7 (alginic acid) 9005-32-7 (Ergosan) 9041-22-9 (beta-glucan) 53238-80-5 (beta-glucan) 9041-22-9 (Macrogard) 53238-80-5 (Macrogard) 9001-63-2 (lysozyme) 9001-63-2 (EC 3.2.1.17)

L188 ANSWER 45 OF 63 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN

ACCESSION NUMBER:

2004:12478 BIOSIS

DOCUMENT NUMBER:

PREV200400016678

TITLE:

Vaccine adjuvants: Role and mechanisms of action in vaccine

immunogenicity.

AUTHOR(S):

Marciani, Dante J. [Reprint Author]

CORPORATE SOURCE:

Galenica Pharmaceuticals, Inc., 2800 Milan Court, Suite

118, Birmingham, AL, 35211, USA

marciani.gpi@att.net

SOURCE:

Drug Discovery Today, (15 October 2003) Vol. 8, No. 20, pp.

934-943. print.

ISSN: 1359-6446 (ISSN print).

DOCUMENT TYPE:

Article

General Review; (Literature Review)

LANGUAGE:

English

ENTRY DATE:

Entered STN: 24 Dec 2003

Last Updated on STN: 24 Dec 2003

ABSTRACT: Inactivated vaccines require adjuvants to stimulate an immune
\*\*\*response.\*\*\* The choice of adjuvant or immune enhancer determines whether
the immune response is effective, ineffective or damaging.
Accordingly, there is a need for new adjuvants that stimulate the appropriate
immunity, for example, T cell immunity for intracellular
pathogens and cancer vaccines. In several adjuvants, the identification of
chemical groups that interact with specific cell toll-like receptors (innate
immunity) or receptors for costimulatory ligands (adaptive immunity),
has enabled the establishment of structure-function relationships that are
useful in the design of new adjuvants. Because of the crucial
\*\*\*immunomodulating\*\*\* role of adjuvants, sub-unit vaccine development will

remain dependent on new adjuvants.
CONCEPT CODE: Cytology - Animal

Biochemistry studies - General 10060

02506

Biochemistry studies - Lipids 10066

Biochemistry studies - Carbohydrates 10068

Pathology - Therapy 12512

Blood - Blood and lymph studies 15002 Blood - Blood cell studies 15004

Pharmacology - General 22002

Immunology - General and methods 34502

INDEX TERMS: Major Concepts

Biochemistry and Molecular Biophysics; Immune System (Chemical Coordination and Homeostasis); Pharmaceuticals

(Pharmacology)

INDEX TERMS: Parts, Structures, & Systems of Organisms

T cells: blood and lymphatics,

immune system

INDEX TERMS: Chemicals & Biochemicals

Toll-like receptors; alpha-galactosylceramide: vaccine

adjuvant; bacterial CpG-DNA: vaccine adjuvant;

beta-glucans: vaccine adjuvant; cancer

vaccines: vaccine; co-stimulatory

ligands: vaccine adjuvant; imidazoquinolines: vaccine adjuvant; lipopolysaccharide [endotoxin]: vaccine adjuvant; small synthetic compounds: vaccine adjuvant; vaccine adjuvants: classification, mechanisms of action,

vaccine component, pharmaceutical

INDEX TERMS: Miscellaneous Descriptors

T cell immunity; adaptive immunity; immune response stimulation; innate

immunity; intracellular pathogens; structure-function

relationships; vaccine development; vaccine

immunogenicity

ORGANISM: Classifier

Vertebrata 85150

Super Taxa

Chordata; Animalia

Organism Name

vertebrate (common)

Taxa Notes

Animals, Chordates, Nonhuman Vertebrates, Vertebrates

REGISTRY NUMBER: 209533-83-5 (alpha-galactosylceramide)

9041-22-9 (beta-glucans)

L188 ANSWER 46 OF 63 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on

STN

ACCESSION NUMBER: 2004:144156 BIOSIS DOCUMENT NUMBER: PREV200400144017

TITLE: Immunopotentiating properties of lentinan (1

fwdarw 3)-beta-D-glucan extracted from

culinary-medicinal Shiitake mushroom Lentinus edodes

(Berk.) Singer (Agaricomycetideae).

AUTHOR(S): Yap, Ann-Teck; Ng, Mah-Lee [Reprint Author]

CORPORATE SOURCE: Department of Microbiology, National University of

Singapore, 5 Science Drive 2, Singapore, 117597, Singapore

micngml@nus.edu.sg

SOURCE: International Journal of Medicinal Mushrooms, (2003) Vol.

5, No. 4, pp. 339-358. print. ISSN: 1521-9437 (ISSN print).

DOCUMENT TYPE: Article LANGUAGE: English

ENTRY DATE: Entered STN: 10 Mar 2004

Last Updated on STN: 10 Mar 2004

ABSTRACT: The immunopotentiating efficacy of lentinan, a fully purified (1fwdarw3)-beta-D-glucan extracted from culinary-medicinal mushroom Lentinus edodes (Berk.) Singer was investigated both in vitro and in vivo. The oral administration of lentinan to AKR mice exerted strong antitumor activity, resulting in raised levels of lymphocytokines such as IFN-gamma, TNF-alpha, IL-2, and IL-1alpha. cultures of murine macrophages CRL-2019, B-lymphocytes HB-284, and T-\*\*\*lymphocytes\*\*\* CRL-8179, which were treated with lentinan, showed high levels of activation using flow cytometry. Activated immunocytes, particularly the T-helper cells by lentinan, might render the physiological constitutions of the host highly cancer and infection resistant. Adoptive immunotherapy of the immunodeficient mice such as the nude (athymic) mice, B-cell deficient mice, and SCID (severe combined immunodeficient) mice via the transfer of the lentinan-activated immunocytes resulted in the inhibition of tumor growth. Lentinan appeared to represent a unique class of host defense potentiators (HDP), protecting the hosts from the side effects of conventional therapeutic measures and improving various kinds of immunological parameters with no toxic side-effects in animal models. Cytology - Animal CONCEPT CODE: 02506 Biochemistry studies - General 10060 Biochemistry studies - Proteins, peptides and amino acids 10064 Pathology - Therapy 12512 Food technology - General and methods 13502 Blood - Blood and lymph studies Blood - Blood cell studies 15004 Endocrine - General 17002 Pharmacology - Immunological processes and allergy Immunology - General and methods 34502 22018 Plant physiology - Chemical constituents 51522 Pharmacognosy and pharmaceutical botany 54000 Major Concepts INDEX TERMS: Biochemistry and Molecular Biophysics; Foods; Immune System (Chemical Coordination and Homeostasis); Pharmacognosy (Pharmacology) Parts, Structures, & Systems of Organisms INDEX TERMS: B-lymphocytes: blood and lymphatics, immune system; T-lymphocytes: blood and lymphatics, immune system; macrophages: blood and lymphatics, immune system Chemicals & Biochemicals INDEX TERMS: IFN-gamma [interferon-gamma]; IL-1-alpha [interleukin-1-alpha]; IL-2 [interleukin-2]; TNF-alpha [tumor necrosis factor-alpha]; lentinan-(1-3)beta-D-glucan: immunologic-drug, extraction, immunopotentiating properties INDEX TERMS: Methods & Equipment flow cytometry: histology and cytology techniques, laboratory techniques INDEX TERMS: Miscellaneous Descriptors immunological parameters ORGANISM: Classifier Basidiomycetes 15300 Super Taxa Fungi; Plantae Organism Name Lentinus edodes (species) [shiitake mushroom (common)]: culinary mushroom, medicinal mushroom Taxa Notes Fungi, Microorganisms, Nonvascular Plants, Plants

Jones 10/630143 Page 82

ORGANISM: Classifier

Muridae 86375

Super Taxa

Rodentia; Mammalia; Vertebrata; Chordata; Animalia

Organism Name mouse (common)

Taxa Notes

Animals, Chordates, Mammals, Nonhuman Vertebrates,

Nonhuman Mammals, Rodents, Vertebrates

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STN

ACCESSION NUMBER: 2002:175191 BIOSIS DOCUMENT NUMBER: PREV200200175191

TITLE: Immunization strategies to augment oral vaccination with

DNA and viral vectors expressing HIV envelope glycoprotein.

AUTHOR(S): Wierzbicki, Andrzej; Kiszka, Irena; Kaneko, Hiroshi;

Kmieciak, Dariusz; Wasik, Thomas J.; Gzyl, Jaroslaw;

Kaneko, Yutaro; Kozbor, Danuta [Reprint author]

CORPORATE SOURCE: Center for Neurovirology and Cancer Biology, Temple

University, 1900 North 12th Street, Philadelphia, PA,

19122, USA

dkozbor@astro.temple.edu

SOURCE: Vaccine, (31 January, 2002) Vol. 20, No. 9-10, pp.

1295-1307. print.

CODEN: VACCDE. ISSN: 0264-410X.

DOCUMENT TYPE: Article LANGUAGE: English

ENTRY DATE: Entered STN: 6 Mar 2002

Last Updated on STN: 6 Mar 2002

ABSTRACT: Induction of mucosal immunity to the human immunodeficiency virus (HIV) envelope (env; gp160) glycoprotein has been demonstrated with orally administered recombinant vaccinia virus (rVV) vectors and poly(DL-lactide-coqlycolide) (PLG)-encapsulated plasmid DNA expressing gp160. In this study, we investigated the effect of an oral DNA-prime/rVV-boost vaccine regimen in conjunction with adjuvants on the level of qp160-specific cellular and humoral responses in BALB/c mice. We demonstrated that DNA priming followed by a booster with rVV expressing gp160 (vPE16) significantly augmented env-specific immunity in systemic and mucosal tissues of the immunized mice. Association of vPE16 with liposomes and coadministration of liposome-associated beta -glucan lentinan or IL-2/Ig-encoded plasmid DNA-encapsulated in PLG microparticles triggered the optimal cell-mediated immune (CMI) responses. Lentinan was found to increase env-specific type 1 cytokine production and cytotoxic T-lymphocyte (CTL) activities but had no effect on humoral responses. On the other hand, IL-2/Ig-mediated increases in both type 1 and 2 activities were associated with higher levels of env-specific CTL and antibody responses. Results of these studies demonstrated the effectiveness of oral vaccines with DNA and rVV vectors in conjunction with \*\*\*immunomodulators\*\*\* in inducing specific immune responses in systemic and mucosal tissues.

CONCEPT CODE: Cytology - Animal 02506

Biochemistry studies - Nucleic acids, purines and

pyrimidines 10062

Biochemistry studies - Proteins, peptides and amino acids

10064

Blood - Blood and lymph studies 15002 Blood - Blood cell studies 15004

Endocrine - General 17002

Immunology - General and methods 34502

INDEX TERMS: Major Concepts

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Page 83

Immune System (Chemical Coordination and Homeostasis)

INDEX TERMS: Parts, Structures, & Systems of Organisms

cytotoxic T-lymphocyte: blood and

lymphatics, immune system

INDEX TERMS: Chemicals & Biochemicals

IL-2 [interleukin-2]; glycoprotein 160; human immunodeficiency virus envelope glycoprotein;

immunomodulators; lentinan; plasmid DNA

INDEX TERMS: Methods & Equipment

oral vaccination: prophylactic method

INDEX TERMS: Miscellaneous Descriptors

cell-mediated immune response;

humoral response; immune responses; mucosal immunity

ORGANISM: Classifier

Muridae 86375

Super Taxa

Rodentia; Mammalia; Vertebrata; Chordata; Animalia

Organism Name

mouse Taxa Notes

Animals, Chordates, Mammals, Nonhuman Vertebrates,

Nonhuman Mammals, Rodents, Vertebrates

REGISTRY NUMBER: 37339-90-5 (lentinan)

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STN

ACCESSION NUMBER: 2002:402334 BIOSIS DOCUMENT NUMBER: PREV200200402334

TITLE: Purification, characterization, and modification of

T lymphocyte-stimulating polysaccharide

from spores of Ganoderma lucidum.

AUTHOR(S): Bao, Xing-Feng; Zhen, Yun; Ruan, Li; Fang, Ji-nian [Reprint

author]

CORPORATE SOURCE: Shanghai Institute of Materia Medica, Shanghai Institutes

for Biological Sciences, Chinese Academy of Sciences, 294

Taiyuan Road, Shanghai, 200031, China

jnfang@mail.shcnc.ac.cn

SOURCE: Chemical and Pharmaceutical Bulletin (Tokyo), (May, 2002)

Vol. 50, No. 5, pp. 623-629. print.

CODEN: CPBTAL. ISSN: 0009-2363.

DOCUMENT TYPE: Article LANGUAGE: English

ENTRY DATE: Entered STN: 24 Jul 2002

Last Updated on STN: 24 Jul 2002

ABSTRACT: The hot-water extract of the spores of Ganoderma lucidum was shown to have a stimulating effect on concanavalin A-induced mitogenic activity of

\*\*\*T\*\*\* lymphocytes. Bioassay-guided separation led to the

isolation of a polysaccharide with potent T lymphocyte

-stimulating activity by ethanol fractionation, anion-exchange, and size-exclusion chromatography. Based on the composition and methylation analyses, periodate oxidation, Smith degradation, and NMR spectroscopy, the

native polysaccharide was shown to be a beta-D-(1fwdarw3)-

\*\*\*glucan\*\*\* with branches of terminal glucosyl residues substituted at C-6 of the **glucose** residues in the main chain. The branching ratio is

approximately 20%. A series of sulfated or carboxymethylated derivatives were prepared and their structural features were elucidated by chemical and spectral analyses. The solution conformation and T lymphocyte

proliferation effect of the glucans before and after derivatization were compared and discussed. The data obtained indicate that the introduction of ionic groups would significantly affect the original conformation of the native glucan in aqueous solution and further affect **T lymphocyte** -stimulating activity. The triple-helical structure of the glucans, the nature of the ionic groups, and the density of negative charge were considered to be closely related to this activity.

CONCEPT CODE: Cytology - Animal 02506

Pathology - Therapy 12512

Blood - Blood and lymph studies 15002

Blood - Blood cell studies 15004

Pharmacology - Immunological processes and allergy 22018

Immunology - General and methods 34502

Pharmacognosy and pharmaceutical botany 54000

INDEX TERMS: Major Concepts

Immune System (Chemical Coordination and Homeostasis);

Pharmacognosy (Pharmacology)

INDEX TERMS: Parts, Structures, & Systems of Organisms

T lymphocyte: blood and lymphatics, immune system, proliferation; spores

INDEX TERMS: Chemicals & Biochemicals

beta-D-(1-3)-glucan:

immunologic-drug, immunostimulant-drug,

carboxymethylated derivatives, characterization, modification, purification, sulfated derivatives

ORGANISM: Classifier

Basidiomycetes 15300

Super Taxa

Fungi; Plantae Organism Name

Ganoderma lucidum: medicinal plant

Taxa Notes

Fungi, Microorganisms, Nonvascular Plants, Plants

ORGANISM: Classifier

Muridae 86375

Super Taxa

Rodentia; Mammalia; Vertebrata; Chordata; Animalia

Organism Name

mouse: animal model

Taxa Notes

Animals, Chordates, Mammals, Nonhuman Vertebrates,

Nonhuman Mammals, Rodents, Vertebrates

L188 ANSWER 49 OF 63 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on

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ACCESSION NUMBER: 2003:49690 BIOSIS DOCUMENT NUMBER: PREV200300049690

TITLE: Medicinal mushrooms as a source of antitumor and

immunomodulating polysaccharides.

AUTHOR(S): Wasser, S. P. [Reprint Author]

CORPORATE SOURCE: Institute of Evolution, University of Haifa, Mt. Carmel,

Haifa, 31905, Israel

spwasser@research.haifa.ac.il

SOURCE: Applied Microbiology and Biotechnology, (November 2002)

Vol. 60, No. 3, pp. 258-274. print.

CODEN: AMBIDG. ISSN: 0175-7598.

DOCUMENT TYPE: Article

General Review; (Literature Review)

LANGUAGE: English

ENTRY DATE: Entered STN: 15 Jan 2003

Last Updated on STN: 15 Jan 2003

ABSTRACT: The number of mushrooms on Earth—is estimated at 140,000, yet maybe only 10% (approximately 14,000 named species) are known. Mushrooms comprise a

vast and yet largely untapped source of powerful new pharmaceutical products. In particular, and most importantly for modern medicine, they represent an unlimited source of polysaccharides with antitumor and
\*\*\*immunostimulating\*\*\* properties. Many, if not all, Basidiomycetes
mushrooms contain biologically active polysaccharides in fruit bodies, cultured mycelium, culture broth. Data on mushroom polysaccharides have been collected from 651 species and 7 infraspecific taxa from 182 genera of higher Hetero- and Homobasidiomycetes. These polysaccharides are of different chemical composition, with most belonging to the group of beta-glucans ; these have beta-(1fwdarw3) linkages in the main chain of the and additional beta-(1fwdarw6) branch points that are needed for their antitumor action. High molecular weight glucans appear to be more effective than those of low molecular weight. Chemical modification is often carried out to improve the antitumor activity of polysaccharides and their clinical qualities (mostly water solubility). The main procedures used for chemical improvement are: Smith degradation (oxydo-reducto-hydrolysis), formolysis, and carboxymethylation. Most of the clinical evidence for antitumor activity comes from the commercial polysaccharides lentinan, PSK (krestin), and schizophyllan, but polysaccharides of some other promising medicinal mushroom species also show good results. Their activity is especially beneficial in clinics when used in conjunction with chemotherapy. Mushroom polysaccharides prevent oncogenesis, show direct antitumor activity against various allogeneic and syngeneic tumors, and prevent tumor metastasis. Polysaccharides from mushrooms do not attack cancer cells directly, but produce their antitumor effects by activating different immune responses in the host. The antitumor action of polysaccharides requires an intact Tcomponent; their activity is mediated through a thymus-dependent immune mechanism. Practical application is dependent not only on biological properties, but also on biotechnological availability. The present review analyzes the pecularities of polysaccharides derived from fruiting bodies and cultured mycelium (the two main methods of biotechnological production today) in selected examples of medicinal mushrooms. CONCEPT CODE: Cytology - Animal 02506 Cytology - Human 02508 Biochemistry studies - General 10060 Biochemistry studies - Carbohydrates Blood - Blood and lymph studies 15002 Blood - Blood cell studies 15004 Reproductive system - Physiology and biochemistry Neoplasms - Immunology 24003 Neoplasms - Pathology, clinical aspects and systemic effects 24004 Immunology - General and methods Immunology - Immunopathology, tissue immunology 34508 Plant physiology - Reproduction 51512 Plant physiology - Chemical constituents 51522 Pharmacognosy and pharmaceutical botany INDEX TERMS: Major Concepts Biochemistry and Molecular Biophysics; Immune System (Chemical Coordination and Homeostasis); Pharmacognosy (Pharmacology); Tumor Biology INDEX TERMS: Parts, Structures, & Systems of Organisms T-cells: blood and lymphatics, immune system; cancer cells, inhibition studies; fruit body: reproductive system; macrophages: blood and lymphatics, immune system; mycelium INDEX TERMS: Chemicals & Biochemicals glucans: molecular weights; pharmaceutical products: applications, preparation, sources; polysaccharides: analysis, antitumor properties, biological properties,

chemical composition studies, immunomodulating properties, isolation, pharmacological properties,

sources

INDEX TERMS: Miscellaneous Descriptors

biotechnology; cancer therapeutics; oncogenesis

ORGANISM: Classifier

Animalia 33000

Super Taxa
Animalia
Organism Name

animal (common)

Taxa Notes
Animals

ORGANISM: Classifier

Basidiomycetes 15300

Super Taxa

Fungi; Plantae Organism Name

basidiomycete (common)

mushroom (common): biochemical constituents, medical

Taxa Notes

Fungi, Microorganisms, Nonvascular Plants, Plants

ORGANISM: Classifier

Hominidae 86215

Super Taxa

Primates; Mammalia; Vertebrata; Chordata; Animalia

Organism Name human (common)

Taxa Notes

Animals, Chordates, Humans, Mammals, Primates,

Vertebrates

ORGANISM: Classifier

Muridae 86375

Super Taxa

Rodentia; Mammalia; Vertebrata; Chordata; Animalia

Organism Name mouse (common)

Taxa Notes

Animals, Chordates, Mammals, Nonhuman Vertebrates,

Nonhuman Mammals, Rodents, Vertebrates

REGISTRY NUMBER: 9012-72-0 (glucans)

L188 ANSWER 50 OF 63 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on

stn

ACCESSION NUMBER: 2001:504787 BIOSIS DOCUMENT NUMBER: PREV200100504787

TITLE: Immunomodulating properties of the chitin

-glucanic preparation in vitro.

AUTHOR(S): Nakonechna, A. [Reprint author]; Drannik, G. [Reprint

author]; Gorovoy, L.; Kushko, L. [Reprint author]

CORPORATE SOURCE: Clinical Immunology and Allergology, National Medical

University, Kiev, Ukraine

SOURCE: Allergy (Copenhagen), j(2001) Vdl. 56, No. Supplement 68,

pp. 109. print.

Meeting Info.: XXth Congress of the European Academy of Allergology and Clinical Immunology. Berlin, Germany. May

09-13, 2001.

CODEN: LLRGDY. ISSN: 0105-4538.

DOCUMENT TYPE: Conference; (Meeting)

Conference; Abstract; (Meeting Abstract)

Jones 10/630143

Page 87

LANGUAGE:

English

ENTRY DATE:

Entered STN: 31 Oct 2001

Last Updated on STN: 23 Feb 2002

CONCEPT CODE:

General biology - Symposia, transactions and proceedings

00520

Cytology - Animal 02506 Cytology - Human 02508

Biochemistry studies - Proteins, peptides and amino acids

Blood - Blood and lymph studies 15002 Blood - Blood cell studies 15004 Immunology - General and methods 34502

Major Concepts

Immune System (Chemical Coordination and Homeostasis)

INDEX TERMS:

INDEX TERMS:

Parts, Structures, & Systems of Organisms T lymphocytes: blood and lymphatics,

immune system; immune system: immune system; peripheral blood: blood and lymphatics; phagocytes: immune system

INDEX TERMS:

Chemicals & Biochemicals

CD19 positive antibodies; CD3 positive antibodies; CD4

positive antibodies; CD8 positive antibodies;

beta-1,3-glucan; beta-1,6-

glucan; chitin-glucanic preparation:

immunomodulating properties; interleukin-1

INDEX TERMS:

Miscellaneous Descriptors

Meeting Abstract

ORGANISM:

Classifier

Hominidae 86215

Super Taxa

Primates; Mammalia; Vertebrata; Chordata; Animalia

Organism Name human Taxa Notes

Animals, Chordates, Humans, Mammals, Primates,

Vertebrates

37361-00-5 (beta-1,6-glucan) REGISTRY NUMBER:

L188 ANSWER 51 OF 63 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on

STN

ACCESSION NUMBER:

1996:77116 BIOSIS

DOCUMENT NUMBER:

PREV199698649251

TITLE:

(1 fwdarw 3) -beta-D-glucans as

biological response modifiers: A review of structure-functional activity relationships.

AUTHOR(S):

Bohn, John A.; Bemiller, James N. [Reprint author]

CORPORATE SOURCE:

Whistler Cent. Carbohydrate Res., 1160 Smith Hall, Purdue

Univ., West Lafayette, IN 47907, USA

SOURCE:

Carbohydrate Polymers, (1995) Vol. 28, No. 1, pp. 3-14.

CODEN: CAPOD8. ISSN: 0144-8617.

DOCUMENT TYPE:

Article

General Review; (Literature Review)

LANGUAGE:

English

ENTRY DATE:

Entered STN: 27 Feb 1996

Last Updated on STN: 28 Feb 1996

ABSTRACT: (1 fwdarw 3)-beta-D-Glucans that have B-

\*\*\*beta\*\*\* -qlucopyranosyl units attached by (1 fwdarw 6) linkages as single unit branches enhance the immune system systemically. This enhancement results in antitumor, antibacterial, antiviral, anticoagulatory and wound healing

activities. The (1 fwdarw 3)-beta-D-glucan backbone is

essential. The most active polymers have degrees of branching (DB) between

0.20 and 0.33. Data suggest both that triple helical structures formed from high molecular weight polymers are possibly important for \*\*\*immunopotentiating\*\*\* activity and that activity is independent of any specific ordered structure. Other data indicate that it is the distribution of the branch units along the backbone chain that is responsible for activity. There are data that indicate both that beta-D-glucopyranosyl units are required for immunopotentiating activity and that the specific nature of the substituent is unimportant. There are also data that indicate both that the more water-soluble polymers are more active (up to a certain degree of substitution (DS) or DB) and that some insoluble aggregates are more stimulatory than the soluble polymers. The best conclusion at this time is that the immunopotentiating activity of (1 fwdarw 3)-beta -D- glucans depends on a helical conformation and on the presence of hydrophilic groups located on the outside surface of the helix. \*\*\*Immunopotentiation\*\*\* effected by binding of a (1 fwdarw 3)-beta -glucan molecule or particle probably includes activation of cytotoxic macrophages, helper T cells, and NK cells, promotion of T cell differentiation, and activation of the alternative complement pathway. CONCEPT CODE: Cytology - Animal 02506 Biochemistry studies - Carbohydrates Biophysics - Molecular properties and macromolecules 10506 Anatomy and Histology - Regeneration and transplantation 11107 Blood - Blood and lymph studies Blood - Blood cell studies 15004 Blood - Lymphatic tissue and reticuloendothelial system 15008 Pharmacology - Blood and hematopoietic agents 22008 Neoplasms - Immunology 24003 Neoplasms - Therapeutic agents and therapy Development and Embryology - Morphogenesis 25508 Immunology - Immunopathology, tissue immunology Medical and clinical microbiology - Bacteriology Medical and clinical microbiology - Virology Chemotherapy - Antibacterial agents Chemotherapy - Antiviral agents 38 38504 Major Concepts INDEX TERMS: Biochemistry and Molecular Biophysics; Blood and Lymphatics (Transport and Circulation); Cell Biology; Development; Immune System (Chemical Coordination and Homeostasis); Pharmacology; Physiology; Tumor Biology Chemicals & Biochemicals INDEX TERMS: (1fwdarw3) -BETA-D-GLUCANS Miscellaneous Descriptors INDEX TERMS: ALTERNATIVE COMPLEMENT PATHWAY ACTIVATION; ANTIBACTERIAL ACTIVITY; ANTICOAGULATORY ACTIVITY; ANTITUMOR ACTIVITY; ANTIVIRAL ACTIVITY; CYTOTOXIC MACROPHAGE ACTIVATION; HELPER T CELL; IMMUNE SYSTEM ENHANCEMENT; IMMUNOPOTENTIATION; NATURAL KILLER CELL; T CELL DIFFERENTIATION PROMOTION; WOUND HEALING ACTIVITY Classifier ORGANISM: 33000 Animalia Super Taxa Animalia Organism Name Animalia

Taxa Notes

Jones 10/630143 Page 89

(1991)

Vol. 49, No. 4,

Animals

9051-97-2 ((lfwdarw3)-BETA-D-REGISTRY NUMBER:

GLUCANS)

L188 ANSWER 52 OF 63 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on

STN

1991:504004 BIOSIS ACCESSION NUMBER:

PREV199192126964; BA92:126964 DOCUMENT NUMBER:

AUGMENTATIVE EFFECT OF POLYSACCHARIDE ON TITLE:

IMMUNOMODULATION IN PATIENTS WITH EARLY GASTRIC

CANCER THE EFFECT OF INTERFERON-GAMMA PRODUCT ABILITY.

URATA Y [Reprint author]; KUSAMA M AUTHOR (S):

DEP SURG, TOKYO MED COLL, JPN CORPORATE SOURCE:

SOURCE: Journal of Tokyo Medical College,

pp. 517-529. CODEN: TIDZAH. ISSN: 0040-8905.

DOCUMENT TYPE: Article

FILE SEGMENT: BA

JAPANESE LANGUAGE:

Entered STN: 12 Nov 1991 ENTRY DATE:

Last Updated on STN: 13 Nov 1991

ABSTRACT: It was well known that some of polysaccharides had possessed an

ability of anti-tumor effect.  $\beta$  -1, 3-glucan

(LentinanR) that a typical reagent reside in the action of host defense-surveillance mechanisms against cancer was tested for its capacity to

modulate the non-specific immune responses of lymphocytes and the specific immune responses to regional lymph nodes and peripheral blood lymphocytes in the patients with early gastric cancer. β -qlucan was

administered to patients intravenously: (group 1): 2 mg to 14 patients one week before operation, (group 2): 2mg to 26 patients 2 weeks after operation and thereafter 4 mg for every 2 weeks. As a control group 13 gastric cancer patients without  $\beta$  -glucan and 15 non-cancer patients

were compared. Lymphocyte count, lymphocyte subpopulation (T

\*\*\*cell\*\*\* , B cell), PHA stimulating blastformation test, single and two color flow cytometric analysis that tested as the non-specific immune

had tendency of the augmentative effect, but not significant. \*\*\*response\*\*\* Interferon- $\gamma$  production in fresh lymphocytes of regional lymph nodes and

peripheral blood that were tested as specific immuno response were significantly augmented by pre- and post-operative administration of .

\*\*\*beta.\*\*\* -glucan. These augmenting effects were not dose dependent. Thus, it is likely that some of polysaccharides might be a multiple

cytokine inducer with IFN-y producing ability for early gastric cancer patients. These results suggest that some of the polysaccharides are feasible as an immuno-modulator.

CONCEPT CODE:

Cytology - Human 02508

Biochemistry studies - Proteins, peptides and amino acids

Biochemistry studies - Carbohydrates · 10068

Pathology - Therapy 12512

Digestive system - General and methods 14001

Digestive system - Pathology Blood - Blood cell studies 15004

Blood - Lymphatic tissue and reticuloendothelial system

15008

Endocrine - General 17002

Pharmacology - Clinical pharmacology

Pharmacology - Digestive system

Neoplasms - Immunology 24003

Neoplasms - Pathology, clinical aspects and systemic

effects 24004 Jones 10/630143 Page 90

Neoplasms - Therapeutic agents and therapy 24008

Immunology - General and methods 34502

INDEX TERMS: Major Concepts

Blood and Lymphatics (Transport and Circulation);

Endocrine System (Chemical Coordination and

Homeostasis); Gastroenterology (Human Medicine, Medical Sciences); Oncology (Human Medicine, Medical Sciences);

Pharmacology

INDEX TERMS: Miscellaneous Descriptors

HUMAN BETA GLUCAN

ANTINEOPLASTIC-DRUG LYMPHOCYTE RESPONSE IMMUNOTHERAPY

ORGANISM: Classifier

Hominidae 86215

Super Taxa

Primates; Mammalia; Vertebrata; Chordata; Animalia

Taxa Notes

Animals, Chordates, Humans, Mammals, Primates,

Vertebrates

REGISTRY NUMBER: 9041-22-9 (BETA-GLUCAN)

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ACCESSION NUMBER: 2004053653 EMBASE

TITLE: Modulation of immune response through nutraceutical

interventions: Implications for canine and feline health.

AUTHOR: Hayek M.G.; Massimino S.P.; Ceddia M.A.

CORPORATE SOURCE: M.G. Hayek, The Iams Company Research and Devmt., PO Box

189, Lewisburg, OH 45338, United States. hayek.mg@pg.com

SOURCE: Veterinary Clinics of North America - Small Animal

Practice, (2004) Vol. 34, No. 1, pp. 229-247. .

Refs: 137

ISSN: 0195-5616 CODEN: VCNAA6

PUBLISHER IDENT.: S 0195-5616(03)00126-8

COUNTRY: Un:

United States

DOCUMENT TYPE: Journal; General Review

FILE SEGMENT: 026 Immunology, Serology and Transplantation

029 Clinical Biochemistry

030 Pharmacology

037 Drug Literature Index

LANGUAGE: English SUMMARY LANGUAGE: English

ENTRY DATE: Entered STN: 20040212

Last Updated on STN: 20040212

ABSTRACT: Mounting research demonstrates that certain nutraceutical compounds interact with the immune system. These interactions may be positive or negative depending on the compound or dose administered to the individual. Understanding the mechanisms by which these compounds work should provide opportunities to design nutritional interventions to bolster the health of dogs and cats.

CONTROLLED TERM: Medical Descriptors:

\*immunomodulation

\*immune response

\*diet supplementation

cat

animal health
medical research
immune system
dose response

molecular mechanics

```
animal food
B lymphocyte
macrophage
  helper cell
cell activity
Echinacea
garlic
Ginkgo biloba
ginseng
tea
mushroom
exercise
immunity
drug mechanism
inflammation: DT, drug therapy
antineoplastic activity
autoimmune disease: DT, drug therapy
dog
nonhuman
review
Drug Descriptors:
CD4 antigen: EC, endogenous compound
CD8 antigen: EC, endogenous compound
cytokine: EC, endogenous compound
immunoglobulin G: EC, endogenous compound
immunoglobulin A: EC, endogenous compound
immunoglobulin M: EC, endogenous compound
immunoglobulin D: EC, endogenous compound
immunoglobulin E: EC, endogenous compound
tumor necrosis factor: EC, endogenous compound
gamma interferon: EC, endogenous compound
thiamine: CM, drug comparison
thiamine: DT, drug therapy
thiamine: PD, pharmacology
pyridoxine: CM, drug comparison
pyridoxine: DT, drug therapy
pyridoxine: PD, pharmacology
ascorbic acid: CM, drug comparison
ascorbic acid: DT, drug therapy
ascorbic acid: PD, pharmacology
alpha tocopherol: CM, drug comparison
alpha tocopherol: DT, drug therapy
alpha tocopherol: PD, pharmacology
beta carotene: CM, drug comparison
beta carotene: DT, drug therapy
beta carotene: PD, pharmacology
zinc: CM, drug comparison
zinc: DT, drug therapy
zinc: PD, pharmacology
selenium: CM, drug comparison
selenium: DT, drug therapy
selenium: PD, pharmacology
chromium: CM, drug comparison
chromium: DT, drug therapy
chromium: PD, pharmacology
cysteine: CM, drug comparison
cysteine: TO, drug toxicity
cysteine: PD, pharmacology
glutamine: CM, drug comparison
glutamine: DT, drug therapy
```

Jones

10/630143 Page 92 Jones

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glutamine: PD, pharmacology
                    selenomethionine: CM, drug comparison
                    selenomethionine: DT, drug therapy
                    selenomethionine: PD, pharmacology
                    garlic extract: CM, drug comparison
                    garlic extract: DT, drug therapy
                    garlic extract: PD, pharmacology
                    green tea extract: CM, drug comparison
                    green tea extract: DT, drug therapy
                    green tea extract: PD, pharmacology
                    isoflavone derivative: PD, pharmacology
                      beta glucan: DT, drug therapy
                    beta glucan: TO, drug toxicity
                      beta glucan: PD, pharmacology
                    Echinacea extract: CM, drug comparison
                    Echinacea extract: DT, drug therapy
                    Echinacea extract: PD, pharmacology
                    genistein: CM, drug comparison
                    genistein: DT, drug therapy
                    genistein: PD, pharmacology
                    Ginkgo biloba extract: CM, drug comparison
                    Ginkgo biloba extract: DT, drug therapy
                    Ginkgo biloba extract: PD, pharmacology
                    ginseng extract: CM, drug comparison
                    ginseng extract: DT, drug therapy
                    ginseng extract: PD, pharmacology
                    unindexed drug
                    (immunoglobulin G) 97794-27-9; (immunoglobulin M)
                    9007-85-6; (immunoglobulin E) 37341-29-0; (gamma
                    interferon) 82115-62-6; (thiamine) 59-43-8, 67-03-8;
                    (pyridoxine) 12001-77-3, 58-56-0, 65-23-6, 8059-24-3;
                    (ascorbic acid) 134-03-2, 15421-15-5, 50-81-7; (alpha
                    tocopherol) 1406-18-4, 1406-70-8, 52225-20-4, 58-95-7,
                    59-02-9; (beta carotene) 7235-40-7; (zinc) 7440-66-6;
                    (selenium) 7782-49-2; (chromium) 16065-83-1, 7440-47-3;
                    (cysteine) 4371-52-2, 52-89-1, 52-90-4; (glutamine)
                    56-85-9, 6899-04-3; (selenomethionine) 1464-42-2,
                    3211-76-5; (genistein) 446-72-0
L188 ANSWER 54 OF 63 EMBASE COPYRIGHT (c) 2006 Elsevier B.V. All rights
     reserved on STN
                    2004219588 EMBASE
                    Chemical and biochemical basis of the potential anti-tumor
                    properties of Ganoderma lucidum.
                    Yeung H.W.; Lu Q.-Y.; Zhang Q.; Go V.L.W.
                    Dr. Q.-Y. Lu, David Geffen School of Medicine, UCLA, 900
                    Veteran Avenue, Los Angeles, CA 90095, United States.
                    qlu@mednet.ucla.edu
                    Current Topics in Nutraceutical Research, (2004) Vol. 2,
                    No. 2, pp. 67-77. .
                    Refs: 55
                    ISSN: 1540-7535 CODEN: CTNRC3
                    United States
                    Journal; General Review
                    016
                            Cancer
                            Public Health, Social Medicine and Epidemiology
                    017
                            Immunology, Serology and Transplantation
                    026
                    029
                            Clinical Biochemistry
                    030
                            Pharmacology
                            Drug Literature Index
                    037
```

CAS REGISTRY NO.:

ACCESSION NUMBER:

CORPORATE SOURCE:

TITLE:

AUTHOR:

SOURCE:

COUNTRY:

DOCUMENT TYPE: FILE SEGMENT:

Jones 10/630143

English LANGUAGE: SUMMARY LANGUAGE: English

Entered STN: 20040604 ENTRY DATE:

Last Updated on STN: 20040604

ABSTRACT: Ganoderma lucidum antitumor substances are divided mainly into alcohol-soluble and water-soluble compounds. Chemical studies on the alcohol-soluble, non-polar compounds resulted in the structure determination of some 20 highly oxygenated lanostanoid-type triterpenes of a diverse chemical nature, including acids, aldehydes and alcohols, which were shown to be cytotoxic against a panel of human and murine tumor cell lines. Preliminary studies on the mechanism of action of some of these cytotoxic triterpenes showed that they inhibited cancer cell growth and reduced Ras oncogene activities. Another group of antitumor compounds, namely polysaccharides, were isolated from hot water extracts of Ganoderma lucidum. The antitumor polysaccharides were found to be  $\beta$ -D-glucans, heteroglycans and peptidoglycans; the structures of some of these polysaccharides have been determined. Mechanistic studies indicated that the antitumor polysaccharides do not have direct cytotoxicity against tumor cells, but activate the host immune system to mount an effective cell-mediated antitumor response. the chemical structures of Ganoderma lucidum antitumor triterpenes and polysaccharides have now been characterized, and should be investigated for their respective mechanisms of action on the carcinogenesis pathway as well as their bioavailability and efficacy in the treatment and prevention of cancer. Copyright .COPYRGT. 2004 by New Century Health Publishers, LLC.

CONTROLLED TERM: Medical Descriptors:

> \*malignant neoplastic disease: DT, drug therapy \*malignant neoplastic disease: PC, prevention

biochemistry

antineoplastic activity

Ganoderma lucidum drug solubility drug structure structure analysis

cytotoxicity

cancer cell culture drug mechanism growth inhibition cancer growth

inhibition kinetics

gene activity drug isolation

immunostimulation

cellular immunity carcinogenesis

drug bioavailability

drug efficacy

sarcoma: DT, drug therapy sarcoma: PC, prevention

liver cell carcinoma: DT, drug therapy liver cell carcinoma: PC, prevention

breast cancer: PC, prevention

T lymphocyte

cytokine production

human

nonhuman

mouse

human cell

animal cell

```
review
                    Drug Descriptors:
                    *Ganoderma lucidum extract: AN, drug analysis
                    *Ganoderma lucidum extract: DT, drug therapy
                    *Ganoderma lucidum extract: PD, pharmacology
                    *Ganoderma lucidum extract: PO, oral drug administration
                    alcohol
                    water
                    triterpene derivative: AN, drug analysis
                    triterpene derivative: DT, drug therapy
                    triterpene derivative: PD, pharmacology
                    triterpene derivative: PO, oral drug administration
                    ganoderic acid: AN, drug analysis
                    qanoderic acid: PD, pharmacology
                    qanodermic acid: AN, drug analysis
                    ganodermic acid: PD, pharmacology
                    lucidenic acid: AN, drug analysis
                    lucidenic acid: PD, pharmacology
                    ganoderiol: AN, drug analysis
                    ganoderiol: PD, pharmacology
                    ganodermnonol: AN, drug analysis
                    ganodermnonol: PD, pharmacology
                    ganodermanondiol: AN, drug analysis
                    ganodermanondiol: PD, pharmacology
                    lucidumol A: AN, drug analysis
                    lucidumol A: PD, pharmacology
                    lucidumol B: AN, drug analysis
                    lucidumol B: PD, pharmacology
                    aldehyde: AN, drug analysis
                    aldehyde: PD, pharmacology
                    ganoderic aldehyde: AN, drug analysis
                    ganoderic aldehyde: PD, pharmacology
                    lucialdehyde: AN, drug analysis
                    lucialdehyde: PD, pharmacology
                    Ras protein: EC, endogenous compound
                    polysaccharide: AN, drug analysis
                    polysaccharide: DT, drug therapy
                    polysaccharide: PD, pharmacology
                    polysaccharide: PO, oral drug administration
                    hot water
                    beta glucan: AN, drug analysis
                      beta glucan: DT, drug therapy
                      beta glucan: PD, pharmacology
                    beta glucan: PO, oral drug administration
                    peptidoglycan polysaccharide: AN, drug analysis
                    peptidoglycan polysaccharide: DT, drug therapy
                    peptidoglycan polysaccharide: PD, pharmacology
                    peptidoglycan polysaccharide: PO, oral drug administration
                    interleukin 1beta: EC, endogenous compound
                    interleukin 6: EC, endogenous compound
                    gamma interferon: EC, endogenous compound
                    tumor necrosis factor alpha: EC, endogenous compound
                    unclassified drug
                    (alcohol) 64-17-5; (water) 7732-18-5; (gamma interferon)
                    82115-62-6
L188 ANSWER 55 OF 63
                     EMBASE COPYRIGHT (c) 2006 Elsevier B.V. All rights
     reserved on STN
                    2003398633 EMBASE
ACCESSION NUMBER:
                    The influence of \beta-glucan on immune responses in
```

CAS REGISTRY NO.:

TITLE:

broiler chicks.

AUTHOR: Guo Y.; Ali R.A.; Qureshi M.A.

CORPORATE SOURCE: M.A. Qureshi, Department of Poultry Science, North Carolina

State University, Raleigh, NC 27695, China.

m qureshi@ncsu.edu

SOURCE: Immunopharmacology and Immunotoxicology, (2003) Vol. 25,

No. 3, pp. 461-472. .

Refs: 27

ISSN: 0892-3973 CODEN: IITOEF

COUNTRY: United States
DOCUMENT TYPE: Journal; Article

FILE SEGMENT: 026 Immunology, Serology and Transplantation

029 Clinical Biochemistry 037 Drug Literature Index

LANGUAGE: English SUMMARY LANGUAGE: English

ENTRY DATE: Entered STN: 20031023

Last Updated on STN: 20031023

ABSTRACT: Beta-1,3/1,6-glucan ( $\beta$ -glucan) was tested as a possible immunomodulator. Chicken macrophages from a macrophage cell line MQ-NCSU and from abdominal exudate of broiler chickens were exposed to various concentrations of  $\beta$ -glucan in vitro. In addition, day-old broiler chicks were fed a diet containing 0, 20, and 40mg/kg  $\beta\text{-glucan}$  in the starter and 0, 20, and 20 mg/kg in the grower diet. Several baseline immune parameters were examined following such exposures. The results showed that  $\beta\text{-glucan}$ exposure increased nitrite and interleukin-1 (IL-1) production as well as induced macrophage to proliferate in culture. However, IL-6 production was not affected. Dietary  $\beta$ -glucan supplementation increased the macrophage phagocytic activity, anti-sheep red blood cells antibody response post-boost, as well as the PHAP-mediated lymphoproliferative response measured as a toe-web swelling. The percentage of CD4(+), CD8(+), and CD4 (+)/CD8(+) double positive lymphocytes in the intestinal intraepithelial leukocytes was increased in  $\beta$ -glucan supplemented chicks. Furthermore, the primary and secondary lymphoid organs such as bursa of Fabricius, thymus and spleen were larger in  $\beta$ -glucan-supplemented chicks as compared to the chicks on basal diet. findings of these studies which showed that  $\beta\text{-glucan}$  improves several baseline immune responses in the chicken imply that  $\beta$ -glucan can be used as a possible immunomodulator in food animals such as the chicken.

CONTROLLED TERM: Medical Descriptors:

\*immunomodulation

animal food immune response

chicken macrophage cell line

concentration response

in vitro study cytokine production diet supplementation sheep erythrocyte phagocytosis

T lymphocyte subpopulation

nonhuman

animal experiment controlled study animal tissue

article

priority journal Drug Descriptors: \*beta glucan: PD, pharmacology

nitrite interleukin 1

CD4 antigen: EC, endogenous compound CD8 antigen: EC, endogenous compound

CAS REGISTRY NO.: (nitrite) 14797-65-0

L188 ANSWER 56 OF 63 EMBASE COPYRIGHT (c) 2006 Elsevier B.V. All rights

reserved on STN

ACCESSION NUMBER: 2003134066 EMBASE

TITLE: Relationship between dendritic cells and the

D-fraction-induced Th-1 dominant response in BALB/c

tumor-bearing mice.

AUTHOR: Harada N.; Kodama N.; Nanba H.

CORPORATE SOURCE: N. Kodama, Department of Microbial Chemistry, Kobe

Pharmaceutical University, 4-19-1, Motoyama-kitamachi, Kobe

658-8558, Japan. n-kodama@kobepharma-u.ac.jp

SOURCE: Cancer Letters, (31 Mar 2003) Vol. 192, No. 2, pp. 181-187.

•

Refs: 21

ISSN: 0304-3835 CODEN: CALEDO

COUNTRY: Ireland

DOCUMENT TYPE: Journal; Article FILE SEGMENT: 016 Cancer

026 Immunology, Serology and Transplantation

030 Pharmacology

037 Drug Literature Index

LANGUAGE: English SUMMARY LANGUAGE: English

ENTRY DATE: Entered STN: 20030410

Last Updated on STN: 20030410

ABSTRACT: Dendritic cells (DCs) are known to not only induce the activation of T cells, but are also associated with the differentiation of T cells. The D-fraction, a  $\beta$ -glucan extracted from maitake (Grifola frondosa) which expresses anti-tumor effects by establishing a helper (Th)-1 dominance in BALB/c mice, enhanced IL-12p70 production by DCs, when the ratio of CD8 $\alpha$ (+) DCs to CD8 $\alpha$ (-) DCs increased. In addition, examination of the tumor rejection effect of D-fraction-stimulated DCs loaded with tumor antigen revealed that tumor growth is inhibited completely by activating CD4(+) T cells and CD8(+) T cells. .COPYRGT. 2003 Elsevier Science Ireland Ltd. All rights reserved.

CONTROLLED TERM: Medical Descriptors:

\*dendritic cell \*tumor rejection

\*Th1 cell

\*cancer inhibition antineoplastic activity

cell stimulation cytotoxic T lymphocyte

cancer graft

cell transplantation cancer cell culture protein synthesis

lymphocyte activation

T lymphocyte nonhuman female mouse

animal experiment

```
Bacteroides fragilis
                    bacterial membrane
                    protein modification
                    protein structure
                    structure activity relation
                    abdominal abscess: DT, drug therapy
                    cellular immunity
                    host
                      T lymphocyte subpopulation
                    antineoplastic activity
                    antimicrobial activity
                    cytokine release
                    Candida albicans
                    Cryptococcus neoformans
                    cell interaction
                    monocyte
                    macrophage
                    neutrophil
                    mushroom
                    cancer: DT, drug therapy
                    human
                    nonhuman
                    clinical trial
                    review
                    Drug Descriptors:
                      *immunomodulating agent: DV, drug development
                      *immunomodulating agent: PD, pharmacology
                    *bacterial polysaccharide: DV, drug development
                    *bacterial polysaccharide: PD, pharmacology
                    *polysaccharide a: DV, drug development
                    *polysaccharide a: PD, pharmacology
                    ampholyte
                    cell adhesion molecule
                    cytokine: EC, endogenous compound
                    krestin: DV, drug development
                    krestin: PD, pharmacology
                    peptidoglycan p: CT, clinical trial
                    peptidoglycan p: DV, drug development
                    peptidoglycan p: DT, drug therapy
                    peptidoglycan p: PD, pharmacology
                    beta 1,3 glucan: DV, drug development
                      beta 1,3 glucan: PD, pharmacology
                    mannan: DV, drug development
                    mannan: PD, pharmacology
                    hyaluronic acid: DV, drug development
                    hyaluronic acid: DT, drug therapy
                    unclassified drug
CAS REGISTRY NO.:
                    (krestin) 66455-27-4; (beta 1,3 glucan) 9051-97-2; (mannan)
                    51395-96-1, 9036-88-8; (hyaluronic acid) 31799-91-4,
                    9004-61-9, 9067-32-7
L188 ANSWER 58 OF 63 EMBASE COPYRIGHT (c) 2006 Elsevier B.V. All rights
     reserved on STN
                    2000023855 EMBASE
ACCESSION NUMBER:
                    Polymeric drugs based on conjugates of synthetic and
                    natural macromolecules. II. Anti-cancer activity of
                    antibody or (Fab')2-targeted conjugates and combined
                    therapy with immunomodulators.
                    Rihova B.; Jelinkova M.; Strohalm J.; Subr V.; Plocova D.;
                    Hovorka O.; Novak M.; Plundrova D.; Germano Y.; Ulbrich K.
```

TITLE:

AUTHOR:

animal model controlled study animal cell article

priority journal Drug Descriptors:

\*beta glucan: PD, pharmacology

\*beta glucan: IP, intraperitoneal drug

administration

tumor antigen: EC, endogenous compound CD8 antigen: EC, endogenous compound interleukin 12: EC, endogenous compound protein p70: EC, endogenous compound CD4 antigen: EC, endogenous compound

CAS REGISTRY NO.: (interleukin 12) 138415-13-1

Yukiguni maitake COMPANY NAME:

L188 ANSWER 57 OF 63 EMBASE COPYRIGHT (c) 2006 Elsevier B.V. All rights

reserved on STN

ACCESSION NUMBER: 2000371581 EMBASE

Polysaccharide immunomodulators as therapeutic agents: TITLE:

Structural aspects and biologic function.

AUTHOR: Tzianabos A.O.

A.O. Tzianabos, Channing Laboratory, Brigham and Women's CORPORATE SOURCE:

Hospital, Harvard Medical School, 181 Longwood Ave.,

Boston, MA 02115, United States.

atzianabos@channing.harvard.edu (2000) Vol. 13, No. 4, pp.

SOURCE: Clinical Microbiology Reviews,

> 523-533. . Refs: 80

ISSN: 0893-8512 CODEN: CMIREX

United States COUNTRY:

DOCUMENT TYPE: Journal; General Review FILE SEGMENT: 004 Microbiology 030 Pharmacology

037 Drug Literature Index

English LANGUAGE: SUMMARY LANGUAGE: English

ENTRY DATE: Entered STN: 20001127

Last Updated on STN: 20001127

ABSTRACT: Polysaccharide immunomodulators were first discovered over 40 years Although very few have been rigorously studied, recent reports have revealed the mechanism of action and structure-function attributes of some of these molecules. Certain polysaccharide immunomodulators have been identified that have profound effects in the regulation of immune responses during the progression of infectious diseases, and studies have begun to define structural aspects of these molecules that govern their function and interaction with cells of the host immune system. These polymers can influence innate and cell-mediated immunity through interactions with T cells, monocytes, macrophages, and polymorphonuclear lymphocytes. The ability to modulate the immune response in an appropriate way can enhance the host's immune response to certain infections. In addition, this strategy can be utilized to augment current treatment regimens such as antimicrobial therapy that are becoming less efficacious with the advent of antibiotic resistance. This review focuses on recent studies that illustrate the structural and biologic activities of specific polysaccharide immunomodulators and outlines their potential for clinical use.

CONTROLLED TERM: Medical Descriptors:

\*immunomodulation

Searched by Barb O'Bryen, STIC 2-2518

Vel. 64, No. 1-3, pp.

CORPORATE SOURCE: B. Rihova, Institute of Microbiology, Academy Sciences

Czech Republic, Videnska 1083, 142 20 Prague 4, Czech

(2000)

Republic

SOURCE: Journal of Controlled Release,

241-261. . Refs: 49

ISSN: 0168-3659 CODEN: JCREEC

PUBLISHER IDENT.:

S 0168-3659(99)00140-6

COUNTRY:

Netherlands

DOCUMENT TYPE:

Journal; Conference Article

FILE SEGMENT:

016 Cancer

O26 Immunology, Serology and Transplantation O27 Biophysics, Bioengineering and Medical

Instrumentation

030 Pharmacology

037 Drug Literature Index

039 Pharmacy 052 Toxicology

LANGUAGE: SUMMARY LANGUAGE:

English English

ENTRY DATE:

Entered STN: 20000202

Last Updated on STN: 20000202

ABSTRACT: We provide data on in vivo targeting of the Thy 1.2 (CDw90) cell surface receptor expressed on neoplastic T cells, mouse EL4 T cell lymphoma. The targeting antibody and the anticancer drug, doxorubicin (DOX) were conjugated to a water-soluble copolymer based on N-(2hydroxypropyl)methacrylamide (HPMA) acting as a carrier responsible for controlled intracellular release of the conjugated drug. The in vivo therapeutic efficacy of HPMA copolymer-bound DOX targeted with anti-EL4 antibody, polyclonal anti-thymocyte globulin (ATG), monoclonal anti-Thy 1.2 antibody or its F(ab')2 fragment was compared with the efficacy of DOX conjugated to HPMA copolymer containing nonspecific IgG or bovine serum albumin (BSA). Anti-EL4 antibody-targeted conjugate caused a significant retardation of tumor growth and an extension of the life span of treated mice. The effect was comparable with that of HPMA copolymer-bound DOX targeted with ATG, anti-Thy 1.2 antibody or its F(ab')2 fragment. However, considerable antitumor effect was seen also in conjugates targeted instead of specific antibodies with syngeneic nonspecific IgG or BSA. Patients with advanced cancer are often immunocompromised due to dysfunction of their immune system induced by cancer and cytotoxic drugs. A significant decrease of unwanted side-effects of targeted drugs against a number of vital organs was already documented. In this study we have compared immunotoxic effects of free DOX with those of its antibody-targeted form on NK cells and cytolytic T lymphocytes (CTLs) isolated from C57BL/10 mice bearing EL4 T cell lymphoma. In the same model we have tested the combination therapy with immunomodulators ( $\beta$ -glucan or AM-2) injected together with targeted daunomycin. We have observed a significant protective effect of targeted DOX against NK cells and CTLs. Moreover, the data revealed that combination therapy considerably enhances antitumor efficacy of the targeted anticancer drug. Copyright (C) 2000 Elsevier Science B.V.

CONTROLLED TERM: Medical Descriptors:

\*antineoplastic activity

drug targeting drug conjugation macromolecule

T cell lymphoma: DT, drug therapy

T lymphocyte

controlled drug release

lifespan

cancer inhibition

cytotoxicity

```
natural killer cell
                      cytotoxic T lymphocyte
                    immunotoxicity
                    cancer cell culture
                    nonhuman
                    male
                    mouse
                    animal experiment
                    animal model
                    controlled study
                    animal cell
                    conference paper
                    priority journal
                    Drug Descriptors:
                    *antibody conjugate: PD, pharmacology
                    *antibody conjugate: PR, pharmaceutics
                    *antibody conjugate: DT, drug therapy
                    *doxorubicin: PD, pharmacology
                    *doxorubicin: PR, pharmaceutics
                    *doxorubicin: TO, drug toxicity
                    *doxorubicin: DT, drug therapy
                    *doxorubicin: IP, intraperitoneal drug administration
                    copolymer: PR, pharmaceutics
                    polymer: PR, pharmaceutics
                      immunomodulating agent: PD, pharmacology
                      immunomodulating agent: DT, drug therapy
                    immunoglobulin F(ab')2 fragment
                    cell surface receptor: EC, endogenous compound
                    n (2 hydroxypropyl) methacrylamide: PR, pharmaceutics
                    drug carrier: PR, pharmaceutics
                    cancer antibody: PR, pharmaceutics
                    thymocyte antibody: PR, pharmaceutics
                    polyclonal antibody: PR, pharmaceutics
                    bovine serum albumin
                    immunoglobulin G
                      beta glucan: PD, pharmacology
                      beta glucan: DT, drug therapy
                    beta glucan: CB, drug combination
                    daunorubicin: PD, pharmacology
                    daunorubicin: DT, drug therapy
                    daunorubicin: CB, drug combination
                    daunorubicin: IP, intraperitoneal drug administration
                    am 2: PD, pharmacology
                    am 2: DT, drug therapy
                    am 2: CB, drug combination
CAS REGISTRY NO.:
                    (doxorubicin) 23214-92-8, 25316-40-9; (n (2
                    hydroxypropyl) methacrylamide) 21442-01-3; (immunoglobulin
                    G) 97794-27-9; (daunorubicin) 12707-28-7, 20830-81-3,
                    23541-50-6
                    (1) Am 2
CHEMICAL NAME:
                    (1) Peregrine pharmaceutical (United States); Farmitalia
COMPANY NAME:
                    Carlo Erba (Italy)
                      EMBASE COPYRIGHT (c) 2006 Elsevier B.V. All rights
L188 ANSWER 59 OF 63
     reserved on STN
                    1999016692 EMBASE
ACCESSION NUMBER:
TITLE:
                    Modulation of endotoxin- and enterotoxin-induced cytokine
                    release by in vivo treatment with \beta-(1,6)-branched
                    \beta-(1,3)-glucan.
```

AUTHOR:

Soltys J.; Quinn M.T.

CORPORATE SOURCE:

M.T. Quinn, Dept. of Veterinary Molecular Biol., Montana

State University, Bozeman, MT 59717, United States.

mquinn@montana.edu

SOURCE:

Infection and Immunity, (1999) Vol. 67, No. 1, pp. 244-252.

Refs: 81

ISSN: 0019-9567 CODEN: INFIBR

COUNTRY:

DOCUMENT TYPE: FILE SEGMENT:

United States Journal; Article Microbiology 004

026 Immunology, Serology and Transplantation

030 Pharmacology

037 Drug Literature Index

LANGUAGE: SUMMARY LANGUAGE: English English

ENTRY DATE:

Entered STN: 19990128

Last Updated on STN: 19990128

ABSTRACT: Leukocytes activated by endotoxin or enterotoxins release proinflammatory cytokines, thereby contributing to the cascade of events leading to septic shock. In the present studies, we analyzed the effects of in vivo administration of a soluble immunomodulator,  $\beta$ -(1,6)-branched  $\beta$ -(1,3)-glucan (soluble β-glucan), on toxin-stimulated cytokine production in monocytes and lymphocytes isolated from treated mice. In vitro stimulation of lymphocytes isolated from soluble  $\beta$ -glucan-treated mice with lipopolysaccharide (LPS) resulted in enhanced production of interleukin-6 (IL-6) and suppressed production of tumor necrosis factor alpha (TNF- $\alpha$ ), while stimulation of these cells with staphylococcal enterotoxin B (SEB) or toxic shock syndrome toxin 1 (TSST-1) resulted in enhanced production of gamma interferon (IFN- $\gamma$ ) and suppressed production of IL-2 and TNF- $\alpha$ compared to that in cells isolated from untreated mice. In vitro stimulation of monocytes isolated from soluble  $\beta$ -glucan-treated mice with LPS also resulted in suppressed TNF- $\alpha$  production, while stimulation of these cells with SEB or TSST-1 resulted in suppressed IL-6 and TNF- $\alpha$  production compared to that in cells isolated from untreated mice. Thus, the overall cytokine pattern of leukocytes from soluble β-glucan-treated mice reflects suppressed production of proinflammatory cytokines, especially TNF- $\alpha$ . Taken together, our results suggest that treatment with soluble  $\beta$ -glucan can modulate the induction cytokines during sepsis, resulting in an overall decrease in host mortality.

CONTROLLED TERM:

Medical Descriptors: \*immunomodulation \*leukocyte activation monocyte

lymphocyte activation

in vivo study in vitro study

sepsis

toxic shock syndrome

nonhuman female mouse

animal experiment animal model controlled study animal cell

intramuscular drug administration

article

priority journal

Drug Descriptors:

\*endotoxin

\*enterotoxin

\*cytokine: EC, endogenous compound

\*beta 1,3 glucan

\*immunomodulating agent

escherichia coli lipopolysaccharide

staphylococcus enterotoxin b

interleukin 6: EC, endogenous compound

tumor necrosis factor alpha: EC, endogenous compound

gamma interferon: EC, endogenous compound interleukin 2: EC, endogenous compound

CAS REGISTRY NO.: (beta 1,3 glucan) 9051-97-2; (staphylococcus enterotoxin b)

39424-53-8; (gamma interferon) 82115-62-6; (interleukin 2)

(1994)

Vol. 17, No.

85898-30-2

COMPANY NAME: Alpha Beta Technology (United States)

L188 ANSWER 60 OF 63 EMBASE COPYRIGHT (c) 2006 Elsevier B.V. All rights

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94181415 EMBASE ACCESSION NUMBER:

DOCUMENT NUMBER: 1994181415

Changes in immune mediators in mouse lung produced by TITLE:

administration of soluble  $(1\rightarrow 3)$ - $\beta$ -D-glucan.

Sakurai T.; Ohno N.; Yadomae T. AUTHOR:

LIMP, Tokyo College of Pharmacy, Horinouchi CORPORATE SOURCE:

1432-1, Hachioji, Tokyo 192-03, Japan

Biological and Pharmaceutical Bulletin, SOURCE:

5, pp. 617-622.

ISSN: 0918-6158 CODEN: BPBLEO

COUNTRY: Japan

DOCUMENT TYPE: Journal; Article

Chest Diseases, Thoracic Surgery and Tuberculosis FILE SEGMENT: 015

Immunology, Serology and Transplantation 026

Pharmacology 030

Drug Literature Index 037

LANGUAGE: English SUMMARY LANGUAGE: English

Entered STN: 940720 ENTRY DATE:

Last Updated on STN: 940720

In this study, we showed that systemic administration of SSG, a ABSTRACT: highly branched soluble  $(1\rightarrow 3)$ - $\beta$ -D-glucan obtained from Sclerotinia sclerotiorum, induced immunological changes in the alveolar space of mice in vivo, assessed by analysing some immune mediators in bronchoalveolar lavage (BAL) fluid. A single i.v. administration of SSG (250 µg/mouse) induced a rapid but transient leakage of the serum components, IgG and fibronectin, into the alveolar space. This was apparent 12 h post-administration and reached a peak on day 2. Similar kinetic changes were found for lysosomal enzyme activities and interferon  $\gamma$  (IFN $\gamma$ ) concentrations in BAL which are markers of activated alveolar macrophages (AMs) or pulmonary T cells. BAL prepared from SSG-treated mice stimulated lysosomal enzyme release from AMs in vitro. However, SSG did not provoke the chronic accumulation of serum proteins in alveoli and did not induce the release of detectable amounts of nitric oxide and the inflammatory cytokines, IL-1, IL-6 and  $TNF\alpha$ , into BAL. However, their mRNAs were detected in lung tissue using the reverse-transcriptase polymerase chain reaction (RT-PCR) technique. Similar results were observed for multiple i.v. administration (250 µg, once a day for 10 consecutive days), and there were a little difference between single and multiple administration. In summary, systemic administration of SSG induces immune responses, including activation of AMs and lymphocytes, but does not provoke chronic inflammation in the alveolar space when administered either as single

or multiple doses. This finding is very important for the clinical application of SSG in immunocompromised hosts as a biological response modifier (BRM) without toxic-side effects on lung tissue.

CONTROLLED TERM: Medical Descriptors:

\*immunomodulation

\*lung

animal experiment animal tissue

article

controlled study

intravenous drug administration

lung lavage

lymphocyte activation
macrophage activation

male mouse nonhuman

polymerase chain reaction

rna synthesis Drug Descriptors:

\*beta 1,3 glucan: PD, pharmacology fibronectin: EC, endogenous compound gamma interferon: EC, endogenous compound immunoglobulin g: EC, endogenous compound

lysozyme: EC, endogenous compound

CAS REGISTRY NO.: (beta 1,3 glucan) 9051-97-2; (fibronectin) 86088-83-7;

(gamma interferon) 82115-62-6; (immunoglobulin g)

97794-27-9; (lysozyme) 9001-63-2

L188 ANSWER 61 OF 63 WPIDS COPYRIGHT 2006 THE THOMSON CORP on STN

ACCESSION NUMBER: 2002-759837 [82] WPIDS

DOC. NO. CPI: C2002-214753

TITLE: New Major Histocompatibility Complex (MHC) molecule

construct, useful for treating, preventing, stabilizing or alleviating a disease involving MHC recognizing cells

e.g., cancer.

DERWENT CLASS: B04 D16

INVENTOR(S): AMELLEM, O; BUUS, S; PETERSEN, L O; RUUD, E; SCHOLLER, J;

WINTHER, L; AAMELLEM, O; RUUB, E; SCHOELLER, J

PATENT ASSIGNEE(S): (DAKO-N) DAKO AS; (DYNA-N) DYNAL BIOTECH ASA; (DAKO-N)

DAKOCYTOMATION DENMARK AS

COUNTRY COUNT: 101

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG MAIN IPC

WO 2002072631 A2 20020919 (200282)\* EN 304 C07K014-705

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ

NL OA PT SD SE SL SZ TR TZ UG ZM ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR

ZW

NO 2003004020 A 20031106 (200380) C07K014-705 EP 1377609 A2 20040107 (200404) EN C07K014-705

R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT

RO SE SI TR

AU 2002240818 A1 20020924 (200433) C07K014-705

Jones 10/630143 Page 104

JP 2005500257 W 20050106 (200505) 439 C07K017-02

### APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 2002072631	A2	WO 2002-DK169	20020313
NO 2003004020	Α	WO 2002-DK169	20020313
		NO 2003-4020	20030911
EP 1377609	A2	EP 2002-706685	20020313
		WO 2002-DK169	20020313
AU 2002240818	A1	AU 2002-240818	20020313
JP 2005500257	W	JP 2002-571544	20020313
		WO 2002-DK169	20020313

### FILING DETAILS:

PATENT NO	KIND	PATENT NO
EP 1377609 AU 2002240818 JP 2005500257	A2 Based on A1 Based on W Based on	WO 2002072631 WO 2002072631 WO 2002072631
PRIORITY APPLN. INFO	US 2001-275470P 2001-435 2001-436 2001-441 2001-275447P 2001-275448P	20010314; DK 20010314; DK 20010314; DK 20010314; US 20010314; US 20010314

## INT. PATENT CLASSIF.:

MAIN: C07K014-705; C07K017-02 A61K038-00; A61K038-17; A61K045-00; A61K047-48; SECONDARY: A61P001-00; A61P001-04; A61P001-16; A61P003-10; A61P007-00; A61P011-00; A61P011-06; A61P013-08; A61P013-10; A61P013-12; A61P015-00; A61P017-00; A61P017-06; A61P019-02; A61P025-00; A61P029-00; A61P031-00; A61P031-12; A61P035-00; A61P035-02; A61P037-02; A61P037-06; A61P037-08; C07K019-00; C12N005-06; C12N013-00; C12Q001-00; C12Q001-04; G01N033-15; G01N033-50; G01N033-53; G01N033-543; G01N033-566; G01N033-58; G01N033-68; G01N037-00

## BASIC ABSTRACT:

WO 200272631 A UPAB: 20021220

NOVELTY - A new Major Histocompatibility Complex (MHC) molecule construct comprising a carrier molecule to which one or more MHC molecules are attached either directly or via one or more entities, is new.

DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for the following:

- (1) detecting the presence or MHC recognizing cells in a sample;
- (2) monitoring MHC recognizing cells;
- (3) establishing a prognosis of a disease involving MHC recognizing cells;
- (4) determining the status of, or the effectiveness of a medicament against, a disease involving MHC recognizing cells;
  - (5) diagnosing a disease involving MHC recognizing cells;
- (6) a therapeutic composition comprising as active ingredient a MHC molecule construct;
- (7) up-regulating, down-regulating or modulating an immune response in an animal, including a human being;
  - (8) treating an animal, including a human being;

```
(9) inducing energy of a cell in animal, including a human being;
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(10) an adoptive cellular immunotherapeutic method;

(11) obtaining MHC recognizing cells; or

(12) producing a therapeutic composition.

ACTIVITY - Cytostatic; Antiinflammatory; Dermatological; Antiasthmatic; Antidiabetic; Anti-HIV; Virucide; Antiarteriosclerotic; Antiulcer; Antirheumatic; Antiarthritic; Antipsoriatic; Immunosuppressive. No biological data is given.

MECHANISM OF ACTION - Gene therapy.

USE - The MHC molecule construct is useful as a therapeutic composition in in vivo or ex vivo therapy, for treating, preventing, stabilizing or alleviating a disease involving MHC recognizing cells, for monitoring MHC recognizing cells or establishing a prognosis of a disease or diagnosing a disease, or determining the status of a disease or the effectiveness of a medicament against a disease, involving MHC recognizing cells, e.g., chronic inflammatory bowel disease such as Crohn's disease or ulcerative colitis, sclerosis, type I diabetes, rheumatoid arthritis, psoriasis, atopic dermatitis, asthma, malignant melanoma, renal carcinoma, breast cancer, lung cancer, cancer of the uterus, cervical cancer, prostate cancer, brain cancer, head and neck cancer, leukemia, cutaneous lymphoma, hepatic carcinoma, colorectal cancer, bladder cancer, rejection-related disease, Graft-versus-host-related disease, or a viral disease associated with hepatitis, Acquired Immunodeficiency Syndrome (AIDS), measles, pox, chicken pox, rubella or herpes. The MHC molecule construct is also useful for flow cytometric, histological or cytological method (all claimed.)

Dwg.0/57

FILE SEGMENT: CPI
FIELD AVAILABILITY: AB; DCN

MANUAL CODES:

AB; DCN
CPI: B02-C01; B02-R; B03-G; B04-B01B; B04-B03C;
B04-B04C7; B04-C01; B04-C02; B04-C03; B04-E01;
B04-F01; B04-G01; B04-H01; B04-H05; B04-K01;
B04-L01; B04-N04; B06-H; B07-H; B11-C07B3; B11-C08E;
B12-K04A; B12-K04B; B14-A02; B14-C09B; B14-E10C;
B14-G02C; B14-H01; B14-K01; B14-N12; B14-N17;
B14-S03; B14-S04; D05-H09; D05-H10; D05-H11;

L188 ANSWER 62 OF 63 WPIDS COPYRIGHT 2006 THE THOMSON CORP on STN

ACCESSION NUMBER: 2002-519746 [55]

DOC. NO. CPI: C2002-147084

TITLE: Novel dietary supplements for use as immunostimulants, containing beta-

glucan and colostrum and/or lactoferrin.

WPIDS

DERWENT CLASS: B04 D13

INVENTOR(S): MCANALLEY, B H

PATENT ASSIGNEE(S): (MCAN-I) MCANALLEY B H; (MANN-N) MANNATECH INC.

COUNTRY COUNT: 97

PATENT INFORMATION:

PATENT NO KIND DATE WEEK LA PG MAIN IPC

WO 2002047612 A2 20020620 (200255) \* EN 34 A61K000-00

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ

NL OA PT SD SE SL SZ TR TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PH PL PT RO

RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG UZ VN YU ZA ZW

US 2002119928 A1 20020829 (200259) A61K038-40

Jones 10/630143 Page 106

AU 2002043267 A 20020624 (200267)

A61K000-00

### APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 2002047612	A2	WO 2001-US45513	20011025
US 2002119928	A1 Provisional	US 2000-244029P	20001027
		US 2001-1439	20011025
AU 2002043267	A	AU 2002-43267	20011025

### FILING DETAILS:

PRIORITY APPLN. INFO: US 2000-244029P 20001027; US

2001-1439 20011025

INT. PATENT CLASSIF.:

MAIN: A61K000-00; A61K038-40 SECONDARY: A61K031-716; A61K031-732

BASIC ABSTRACT:

WO 200247612 A UPAB: 20020829

NOVELTY - Novel dietary supplements containing **beta** - **glucan** and colostrum and/or lactoferrin for supporting and promoting strong immune systems

ACTIVITY - Immunostimulant.

 ${\tt MECHANISM}$  OF ACTION - No specific mechanisms given in source material.

USE - The compositions are useful for supporting and promoting strong immune systems. The compositions are useful for providing a first effect comprising regulation of the immune system, regulation of cytokine release, prevention of autoimmune response from intestinal pathogens, promotion of phagocytosis by neutrophils, stimulation of B cell and antibody secretion, inhibition of mast cell enzyme involved in allergic airway response, enhancement of natural killer cell activity, stimulation of muscle protein synthesis, inhibition of muscle protein breakdown, stimulation of wound healing, stimulation of tissue repair, induction of cartilage formation and bone repair, anti-inflammatory effects, bioregulation during trauma stress, enhancement of hematopoietic activity, increase in insulin-like growth factor in tissues, antidiarrheal effect on gastrointestinal tract infection, stimulation of gastrointestinal tract growth, improvement in function of the gastrointestinal tract, promotion of the growth of beneficial gastrointestinal tract bacteria, lowering blood cholesterol, improving glucose tolerance, reducing average blood glucose in noninsulin dependent diabetics, stimulation of glucose uptake by muscles, inhibition of the binding of bacteria to a host tissue, inhibition of the growth of bacteria, protection against viruses, enhancing activity of antibiotics, antifungal effects, anti-amebic effects, prevention of tumor development, inhibition of tumor cell growth or metastasis, enhancement of natural killer cell toxicity to tumors, improvement in Alzheimer's dementia, antioxidant effects and reaction against bacterial toxins.

The composition comprising beta -glucan and colostrum and/or lactoferrin has a second effect comprising enhancing bile acid excretion, enhancing cholesterol excretion, reducing atherosclerosis, binding heavy metals, stimulation of immune function, resistance to infection, suppression of infection, increase of tissue repair and healing, promotion of body health and athletic performance, promotion of

gastrointestinal tract health, promotion of blood vessel health, promotion of **glucose** utilization and blood sugar balance, improved cancer inhibition, improved metal function and improved toxin related activities (all claimed).

The compositions react with specific cell receptors that cause cells to engulf and destroy bacteria and cellular debris and supplies and enhances natural antibodies. The composition helps regulate the number and activities of circulating immune cells and initiates communication in the immune system which releases chemical messengers to fight infection. The composition supports the immune cell growth and proliferation in the GI tract and binds iron so that it starves bad bacteria, re-routing the iron to be more bio-available for beneficial uses. The composition helps the body remove heavy metals and toxins from cells and help balance the immune system.

ADVANTAGE - The compositions are fast acting, they energize a cascade of immune responses beginning in the mouth and proceeding throughout the body and they optimize the response of natural killer cells B-cells and T-cells which seek out and destroy foreign substances.

Dwg.0/0

FILE SEGMENT: CPI
FIELD AVAILABILITY: AB; DCN

MANUAL CODES: CPI: B04-C02D; B04-N06; B14-A02; B14-A03A; B14-C03;

B14-E02; B14-E10; B14-F03; B14-F06; B14-F07; B14-G03; B14-H01; B14-J01A4; B14-L01; B14-L06; B14-N01; B14-N17B; B14-S04; B14-S08; D03-H01T2

L188 ANSWER 63 OF 63 WPIDS COPYRIGHT 2006 THE THOMSON CORP on STN

ACCESSION NUMBER:

2000-023324 [02] WPIDS

DOC. NO. CPI:

C2001-097584

TITLE:

Polysaccharide adjuvant-antigen conjugates,

useful in pharmaceutical compositions and vaccines to

enhance and potentiate immune responses.

DERWENT CLASS:

B04 C03 D16

INVENTOR(S):

MARCIANI, D J

PATENT ASSIGNEE(S):

(GALE-N) GALENICA PHARM INC

COUNTRY COUNT: 87

PATENT INFORMATION:

PA'	TENT NO	KIND	DATE	WEEK	LA	PG	MAIN IPC
WO	9955715	A2 :	19991104	(200002)	* EN	5	9 C07H003-04

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW NL

OA PT SD SE SL SZ UG ZW

W: AE AL AM AT AU AZ BA BB BG BR BY CA CH CN CU CZ DE DK EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR

TT UA UG US UZ VN YU ZA ZW

AU 9937676 A 19991116 (200015)

1073667 A2 20010207 (200109) EN C07H003-04

R: AT BE CH CY DE DK ES FI FR GB GR IE IT LI LU MC NL PT SE

 JP 2002513028
 W 20020508 (200234)
 67 A61K039-39

 AU 760669
 B 20030522 (200338)
 C07H003-04

 US 6573245
 B1 20030603 (200339)
 A61K031-70

## APPLICATION DETAILS:

PATENT NO	KIND	APPLICATION	DATE
WO 9955715	A2	WO 1999-US9164	19990428
AU 9937676	Α	AU 1999-37676	19990428

ΕP	1073667	A2	ΕP	1999-920096	19990428
			WO	1999-US9164	19990428
JP	2002513028	W	WO	1999-US9164	19990428
			JΡ	2000-545873	19990428
ΑU	760669	В	ΑU	1999-37676	19990428
US	6573245	B1 Provisional	US	1998-83106P	19980428
			US	1999-301115	19990428

#### FILING DETAILS:

PATENT NO	KIND	PATENT NO
AU 9937676 EP 1073667 JP 2002513028	A Based on A2 Based on W Based on	WO 9955715 WO 9955715 WO 9955715
AU 760669	B Previous Publ. Based on	AU 9937676 WO 9955715

PRIORITY APPLN. INFO: US 1998-83106P 19980428; US

1999-301115 19990428

INT. PATENT CLASSIF.:

A61K031-70; A61K039-39; C07H003-04 MAIN:

A61K039-00; A61K039-02; A61K039-04; A61K039-05; SECONDARY: A61K039-07; A61K039-135; A61K039-145; A61K039-165; A61K039-205; A61K039-21; A61K039-23; A61K039-29; A61P031-04; A61P031-10; A61P031-12; A61P031-14; A61P031-16; A61P035-00; A61P037-04; C07H003-06;

C08B037-00; C08B037-06

### BASIC ABSTRACT:

WO 9955715 A UPAB: 20010620

NOVELTY - Polysaccharide adjuvant-antigen conjugates comprising a polysaccharide (I) capable of binding to the surface of antigen presenting cells (APCs) attached to at least 1 molecule with stable carbonyl groups, and at least 1 immunogenic polypeptide or peptide, are new.

DETAILED DESCRIPTION - Polysaccharide adjuvant-antigen conjugates comprising:

- (i) a polysaccharide capable of binding to the surface of antigen presenting cells (APCs);
- (ii) at least 1 molecule with stable carbonyl groups, covalently attached to (I) either directly or via bifunctional linker that keeps the stable carbonyl group intact; and
- (iii) at least 1 polypeptide or peptide capable of eliciting an immunogenic response when covalently attached to (I) either directly or via bifunctional linker.

ACTIVITY - Vaccine; immunogenic; immunopotentiating.

USE - The conjugate may be used in pharmaceutical compositions and vaccines, to enhance and potentiate immune responses in mammals. The conjugate may be used to target and co-stimulate APCs and as vaccine antigens to stimulate T-cell immunity.

ADVANTAGE - The conjugate is stable, easy to reproduce and non-toxic. Dwg.0/0

FILE SEGMENT: CPI

FIELD AVAILABILITY: AB; GI; DCN

MANUAL CODES: CPI: B04-B04C1; B04-C01; B04-C02; B04-K01; B04-N03;

B06-A01; B06-B01; B06-D02; B07-A01; B07-B01; B07-D02; B07-D03; B07-D04C; B07-D10; B07-D12; B07-E01; B10-A06; B10-B01A; B10-B01B; B10-B02J; B10-C04D; B10-D01; B10-E02; B14-S11; C04-B04C1; C04-C01; C04-C02; C04-K01; C04-N03; C06-A01;

Searched by Barb O'Bryen, STIC 2-2518

C06-B01; C06-D02; C07-A01; C07-B01; C07-D02; C07-D03; C07-D04C; C07-D10; C07-D12; C07-E01; C10-A06; C10-B01A; C10-B01B; C10-B02J; C10-C04D; C10-D01; C10-E02; C14-S11; D05-H07

FILE 'HOME' ENTERED AT 13:10:34 ON 03 FEB 2006

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Mis Contraction of the Contracti

# => d his nofile

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(FILE 'HOME' ENTERED AT 11:33:41 ON 03 FEB 2006)
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               1 SEA ABB=ON US2003-630143/AP
L1
                 SET NOTICE LOGIN SEARCH
                 SET LINE LOGIN
                 SET DETAIL LOGIN
                 D SCAN
               0 S HUNTER K?/UA
L*** DEL
            363 SEA ABB=ON HUNTER K?/AU
L2
            110 SEA ABB=ON GAULT R?/AU
L3
            339 SEA ABB=ON JORDAN F?/AU
L4
               3 SEA ABB=ON L2 AND L3 AND L4
L5
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L6
               1 SEA ABB=ON CHITIN/CN
                 E N-ACETYLGLUCOSAMINE/CN
               1 SEA ABB=ON N-ACETYLGLUCOSAMINE/CN
L7
                 E B(1,6) - GLUCAN/CN
               1 SEA ABB=ON 37361-00-5
L8
                 D SCAN
                 E B-D-GLUCAN, (1.FWDARW.3)/CN
               2 SEA ABB=ON "B-D-GLUCAN, (1.FWDARW.3)-"/CN
L9
               2 SEA ABB=ON GLUCOSE/CN
L10
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L11
           9870 SEA ABB=ON GLUCAN#/OBI
           2385 SEA ABB=ON L8 OR L9
L12
L13
         185348 SEA ABB=ON
                             L10
           8812 SEA ABB=ON L6
L14
            457 SEA ABB=ON L7/D
L15
                 D SCAN L1
L16
               8 SEA ABB=ON (L2 OR L3 OR L4) AND (L11 OR L12)
                 E T CELL (LYMPHOCYTE) +ALL/CT
                   "T CELL (LYMPHOCYTE) "+OLD/CT
                 E
         117300 SEA ABB=ON T/OBI(L)(CELL# OR LYMPHOCYTE#)/CW
15587 SEA ABB=ON IMMUNOSTIMULANTS/CT
12883 SEA ABB=ON B7#/BI
L17
L18
L19
          18994 SEA ABB=ON ADJUVANT#/OBI
L20
                 E IMMUNITY+ALL/CT
L21
          48414 SEA ABB=ON IMMUNITY/CT
           9283 SEA ABB=ON IMMUNIZATION/CT
L22
          46764 SEA ABB=ON VACCINES/CT
8960 SEA ABB=ON IMMUNOMODULATORS/CT
5 SEA ABB=ON L16 AND ((L17 OR L18 OR L19 OR L20 OR L21 OR L22
L23
L24
L25
                 OR L23 OR L24) OR (L13 OR L14 OR L15))
L26
              53 SEA ABB=ON (L11 OR L12) AND L13 AND (L14 OR L15)
            646 SEA ABB=ON
                             (L11 OR L12) AND (L17 OR L18 OR L19 OR L20 OR L21
L27
                 OR L22 OR L23 OR L24)
L28
               3 SEA ABB=ON L26 AND L27
L29
           1576 SEA ABB=ON (L11 OR L12)(L)(THU OR BAC OR PAC OR PKT OR
                 DMA)/RL
L30
              35 SEA ABB=ON L29 AND L17 AND (L18 OR L19 OR L20 OR L21 OR L22
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OR L23 OR L24)
L31
              3 SEA ABB=ON L30 AND (L13 OR L14 OR L15)
L32
              1 SEA ABB=ON L28 AND L31
                D SCAN L1
            23 SEA ABB=ON L29 AND L17 AND L18
L33
          3489 SEA ABB=ON COSTIMULA?/OBI OR CO STIMULA?/OBI
L34
            13 SEA ABB=ON L29 AND L17 AND L18 AND ((L19 OR L20) OR L34)
L35
              9 SEA ABB=ON L35 NOT (L1 OR L5 OR L25 OR L28 OR L31)
L36
                D SCAN TI
              1 SEA ABB=ON ENHANCING/TI AND L36
L37
                D SCAN
            228 SEA ABB=ON L29(L)(L19 OR L20 OR L34 OR IMMUN?/OBI)
L38
             26 SEA ABB=ON L38 AND L17
L39
             14 SEA ABB=ON L38 AND L17 AND L18
L40
                E T-CELL/CT
                E E3+AL
                E E3+ALL
                E E2+ALL
         100847 SEA ABB=ON LYMPHOCYTE#/CW(L)T/OBI
L41
             12 SEA ABB=ON L38 AND L17 AND L18 AND L41
L42
     FILE 'MEDLINE' ENTERED AT 11:56:25 ON 03 FEB 2006
            383 SEA ABB=ON HUNTER K?/AU
L43
            19 SEA ABB=ON GAULT R?/AU
L44
            301 SEA ABB=ON JORDAN F?/AU
L45
              2 SEA ABB=ON L44 AND (L43 OR L45)
L46
                D TRIAL 1-2
          87690 SEA ABB=ON GLUCANS+NT/CT
L47
          22108 SEA ABB=ON ADJUVANTS, IMMUNOLOGIC/CT
L48
                E IMMUNOSTIM/CT
                E E4+ALL
              1 SEA ABB=ON (L43 OR L44 OR L45) AND L47 AND L48
L49
              8 SEA ABB=ON (L43 OR L44 OR L45) AND L47
L50
                D TRIAL 1-8
           3885 SEA ABB=ON BETA-GLUCANS/CT OR GLUCANS/CT
4 SEA ABB=ON (L43 OR L44 OR L45) AND L51
1137 SEA ABB=ON L51(L)(IM OR ME)/CT
L51
L52
L53
             46 SEA ABB=ON L53 AND L48
L54
                E T-CELL/CT
                E E3+ALL
         176666 SEA ABB=ON T-LYMPHOCYTES+NT/CT
L55
              1 SEA ABB=ON L54 AND L55
L56
             16 SEA ABB=ON L53/MAJ AND L48/MAJ
L57
                D TRIAL 1-16
L58
          79475 SEA ABB=ON LYMPHOCYTE ACTIVATION/CT
          34369 SEA ABB=ON IMMUNIZATION/CT
L59
          7525 SEA ABB=ON VACCINES/CT
L60
          93143 SEA ABB=ON GLUCOSE/CT
L61
          3543 SEA ABB=ON CHITIN/CT
L62
                E N-ACETYLGLUCOS/CT
           9797 SEA ABB=ON GLUCOSAMINE+NT/CT
L63
              8 SEA ABB=ON L51 AND L61 AND (L62 OR L63)
L64
                D TRIAL 1-8
              4 SEA ABB=ON L54 AND L58 AND L48
L65
     FILE 'EMBASE' ENTERED AT 12:10:44 ON 03 FEB 2006
            287 SEA ABB=ON HUNTER K?/AU
L66
            15 SEA ABB=ON GAULT R?/AU
L67
            218 SEA ABB=ON JORDAN F?/AU
L68
              0 SEA ABB=ON L67 AND (L66 OR L68)
L69
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E BETA-GLUCAN/CT
                E E4+ALL
                E E2+ALL
L70
            886 SEA ABB=ON BETA GLUCAN/CT
L71
           691 SEA ABB=ON (L8 OR L9)
                D TRIAL 1-5
L72
            470 SEA ABB=ON BETA 1,3 GLUCAN/CT
                E BETA 1,6 GLUCAN/CT
L73
             38 SEA ABB=ON BETA 1,6 GLUCAN/CT
L74
            221 SEA ABB=ON L71 NOT (L72 OR L73)
                D TRIAL 1-5
                D SCAN L1
     FILE 'CAPLUS' ENTERED AT 12:13:38 ON 03 FEB 2006
               D SCAN L1
L*** DEL
              3 S L66-L68 AND L70-L73
     FILE 'EMBASE' ENTERED AT 12:14:58 ON 03 FEB 2006
              1 SEA ABB=ON (L66 OR L67 OR L68) AND (L70 OR L71 OR L72 OR L73)
L75
                D TRIAL
                E IMMUNOPOTEN/CT
                E E8+ALL
          13726 SEA ABB=ON IMMUNOPOTENTIATION+NT/CT
L76
                D BIB L75
                E IMMUNOMOD/CT
L77
           6094 SEA ABB=ON IMMUNOMODULATING AGENT/CT
                E IMMUNOMODULATION/CT
                E E3+ALL
L78
          25681 SEA ABB=ON IMMUNOMODULATION/CT
                E IMMUNOPOTENTIATING/CT
            122 SEA ABB=ON (L70 OR L71 OR L72 OR L73) AND (L76 OR L77 OR L78)
L79
L80
             31 SEA ABB=ON (L70 OR L71 OR L72 OR L73) AND L76
            214 SEA ABB=ON (L70 OR (L72 OR L73))(L)(DT OR PD OR PK OR AD OR
L81
                DO)/CT
             13 SEA ABB=ON L81 AND L76
L82
                E T-CELL/CT
                E E9+ALL
         165379 SEA ABB=ON T LYMPHOCYTE+NT/CT
L83
              2 SEA ABB=ON L81 AND L76 AND L83
L84
             14 SEA ABB=ON (L70 OR L71 OR L72 OR L73) AND (L76 OR L77 OR L78)
L85
                AND L83
                D TRIAL 1-5
L86
              8 SEA ABB=ON L81 AND (L76 OR L77 OR L78) AND L83
         102450 SEA ABB=ON GLUCOSE/CT
L87
           1783 SEA ABB=ON CHITIN/CT
L88
                E N ACETYLGLUC/CT
                E N ACETYLGLUCOSAMINE DE/CT
           2922 SEA ABB=ON N ACETYLGLUCOSAMINE/CT OR N ACETYLGLUCOSAMINE
L89
                DERIVATIVE/CT
L90
             10 SEA ABB=ON (L70 OR L71 OR L72 OR L73) AND L87 AND (L88 OR
                L89)
                D TRIAL 1-10
L91
            741 SEA ABB=ON L70/MAJ OR L72/MAJ OR L73/MAJ
                E LYMPHOCYTE ACTIVATION+ALL/CT
L92
          11887 SEA ABB=ON LYMPHOCYTE ACTIVATION/CT
              4 SEA ABB=ON L91 AND L92
L93
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D TRIAL 1-4

```
FILE 'DRUGU' ENTERED AT 12:27:34 ON 03 FEB 2006
              44 SEA ABB=ON HUNTER K?/AU
L94
               2 SEA ABB=ON GAULT R?/AU
L95
               7 SEA ABB=ON JORDAN F?/AU
L96
               1 SEA ABB=ON (L95 OR L96) AND L94
L97
                 D TRIAL
                 E BETA GLUCAN/CT
                 E BETA-GLUCAN/CT
L98
              29 SEA ABB=ON (L8 OR L9)
                 D TRIAL 1-5
                 E GLUCAN-BETA-1,3-/CT
              80 SEA ABB=ON GLUCAN-BETA-1,3/CT OR GLUCAN-BETA-1,3-D/CT
L99
                 E GLUCAN-BETA-1,6-/CT
L100
               2 SEA ABB=ON GLUCAN-BETA-1,6-D/CT
                 E GLUCAN-BETA/CT
               4 SEA ABB=ON GLUCAN-BETA/CT
L101
                 E T-LYMPH/CT
                 E E4+ALL
          17383 SEA ABB=ON THYMOCYTE/CT
L102
               O SEA ABB=ON (L94 OR L95 OR L96) AND (L98 OR L99 OR L100 OR
L103
                 L101)
               4 SEA ABB=ON (L98 OR L99 OR L100 OR L101) AND L102
L104
                 D TRIAL 1-4
          35519 SEA ABB=ON IMMUNOSTIMULANT#/CT
10349 SEA ABB=ON IMMUNE-RESPONSE/CT
L105
L106
          32092 SEA ABB=ON LYMPHOCYTE/CT
L107
          61981 SEA ABB=ON BIOLOGICAL RESPONSE MODIFIERS/CC
L108
              11 SEA ABB=ON (L98 OR L99 OR L100 OR L101) AND L107
11 SEA ABB=ON (L98 OR L99 OR L100 OR L101) AND L107 AND (L105 OR
L109
L110
                 L106 OR L107 OR L108)
               8 SEA ABB=ON (L98 OR L99 OR L100 OR L101) AND L107 AND ((L105
L111
                 OR L106) OR L108)
                 D TRIAL 1-4
     FILE 'WPIDS' ENTERED AT 12:36:31 ON 03 FEB 2006
L112
            123 SEA ABB=ON HUNTER K?/AU
              24 SEA ABB=ON GAULT R?/AU
L113
              68 SEA ABB=ON JORDAN F?/AU
L114
               2 SEA ABB=ON L112 AND L113 AND L114
L115
                 D TRIAL 1-2
           2241 SEA ABB=ON GLUCAN#
L116
          9860 SEA ABB=ON IMMUNE RESPONSE
13730 SEA ABB=ON ADJUVANT#
L117
L118
           5852 SEA ABB=ON IMMUNOSTIMULA?
436 SEA ABB=ON IMMUNOPOTENTIAT?
L119
L120
            483 SEA ABB=ON COSTIMULA? OR CO STIMULA?
L121
           2049 SEA ABB=ON IMMUN#(W) (STIMULA? OR POTENTIAT? OR MODULAT?)
L122
           8323 SEA ABB=ON IMMUNOMODULAT?
L123
           1232 SEA ABB=ON B7
L124
          11598 SEA ABB=ON T(W) (CELL# OR LYMPHOCYTE#)
L125
L126
               2 SEA ABB=ON (L112 OR L113 OR L114) AND L116 AND (L117 OR L118
                 OR L119 OR L120 OR L121 OR L122 OR L123 OR L124 OR L125)
              19 SEA ABB=ON L116 AND L125 AND (L117 OR L118 OR L119 OR L120 OR
L127
                 L121 OR L122 OR L123 OR L124)
L128
          35231 SEA ABB=ON GLUCOSE
           4807 SEA ABB=ON CHITIN OR ACETYLGLUCOSAMINE OR ACETYL(W)GLUCOSAMINE
L129
L130
               2 SEA ABB=ON L127 AND L128 AND L129
               7 SEA ABB=ON L127 AND (L128 OR L129)
L131
           1272 SEA ABB=ON BETA(3A)L116
L132
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2 SEA ABB=ON (L141 OR L142) AND L144 AND (L147 AND L150)

L168 L169 L170 L171

L172

L150)

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FILE 'STNGUIDE' ENTERED AT 13:03:05 ON 03 FEB 2006
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FILE 'CAPLUS' ENTERED AT 13:04:31 ON 03 FEB 2006

D QUE L1

D QUE L5

D QUE L25

L173 5 SEA ABB=ON L1 OR L5 OR L25

FILE 'MEDLINE' ENTERED AT 13:04:33 ON 03 FEB 2006

D QUE L46

D QUE L62

L174 5 SEA ABB=ON L46 OR L52

FILE 'EMBASE' ENTERED AT 13:04:34 ON 03 FEB 2006

D QUE L75

D QUE L69

FILE 'DRUGU' ENTERED AT 13:04:35 ON 03 FEB 2006

D QUE L97

D QUE L103

FILE 'WPIDS' ENTERED AT 13:04:37 ON 03 FEB 2006

D QUE L115

D QUE L126

L175 2 SEA ABB=ON L115 OR L126

FILE 'BIOSIS' ENTERED AT 13:04:40 ON 03 FEB 2006

D QUE L140

D QUE L143

L176 6 SEA ABB=ON L140 OR L143

FILE 'STNGUIDE' ENTERED AT 13:05:18 ON 03 FEB 2006

FILE 'CAPLUS' ENTERED AT 13:05:44 ON 03 FEB 2006

D QUE L1

D QUE L5

D QUE L25

L177 5 SEA ABB=ON L1 OR L5 OR L25

FILE 'MEDLINE' ENTERED AT 13:05:46 ON 03 FEB 2006

D QUE L46

D QUE L52

L178 5 SEA ABB=ON L46 OR L52

FILE 'EMBASE' ENTERED AT 13:05:47 ON 03 FEB 2006

D QUE L75

D QUE L69

FILE 'DRUGU' ENTERED AT 13:05:48 ON 03 FEB 2006

D QUE L97

D QUE L103

FILE 'WPIDS' ENTERED AT 13:05:49 ON 03 FEB 2006

D QUE L115

D QUE L126

L179 2 SEA ABB=ON L115 OR L126

FILE 'BIOSIS' ENTERED AT 13:05:52 ON 03 FEB 2006 D QUE L140

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D QUE L143
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L180 6 SEA ABB=ON L140 OR L143

FILE 'STNGUIDE' ENTERED AT 13:05:54 ON 03 FEB 2006

FILE 'MEDLINE, DRUGU, CAPLUS, BIOSIS, EMBASE, WPIDS' ENTERED AT 13:06:33
ON 03 FEB 2006

L181 11 DUP REM L178 L97 L177 L180 L75 L179 (9 DUPLICATES REMOVED)

ANSWERS '1-5' FROM FILE MEDLINE ANSWERS '6-8' FROM FILE CAPLUS ANSWERS '9-11' FROM FILE BIOSIS

D IALL 1-5

D IBIB ED ABS HITIND 6-8

D IALL 9-11

FILE 'STNGUIDE' ENTERED AT 13:07:02 ON 03 FEB 2006

FILE 'CAPLUS' ENTERED AT 13:09:14 ON 03 FEB 2006

D QUE L28

D QUE L31

D QUE L42

L182 12 SEA ABB=ON (L28 OR L31 OR L42) NOT L177

FILE 'MEDLINE' ENTERED AT 13:09:16 ON 03 FEB 2006

D QUE L56

D QUE L57

D QUE L65

L183 18 SEA ABB=ON (L56 OR L57 OR L65) NOT L178

FILE 'EMBASE' ENTERED AT 13:09:18 ON 03 FEB 2006

D QUE L86

D QUE L93

L184 12 SEA ABB=ON (L86 OR L93) NOT L75

FILE 'DRUGU' ENTERED AT 13:09:20 ON 03 FEB 2006

D QUE L111

D QUE L104

L185 10 SEA ABB=ON (L111 OR L104) NOT L97

FILE 'WPIDS' ENTERED AT 13:09:22 ON 03 FEB 2006

D QUE L130

D QUE L134

L186 4 SEA ABB=ON (L130 OR L134) NOT L179

FILE 'BIOSIS' ENTERED AT 13:09:25 ON 03 FEB 2006

D QUE L160

D QUE L163

D QUE L167

D QUE L168

D QUE L170

D QUE L171

D QUE L172

L187 13 SEA ABB=ON (L160 OR L163 OR L167 OR L168 OR (L170 OR L171 OR L172)) NOT L180

FILE 'STNGUIDE' ENTERED AT 13:09:32 ON 03 FEB 2006

FILE 'MEDLINE, DRUGU, CAPLUS, BIOSIS, EMBASE, WPIDS' ENTERED AT 13:09:58 ON 03 FEB 2006

L188 63 DUP REM L183 L185 L182 L187 L184 L186 (6 DUPLICATES REMOVED)

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ANSWERS '1-18' FROM FILE MEDLINE
ANSWERS '19-28' FROM FILE DRUGU
ANSWERS '29-40' FROM FILE CAPLUS
ANSWERS '41-52' FROM FILE BIOSIS
ANSWERS '53-60' FROM FILE EMBASE
ANSWERS '61-63' FROM FILE WPIDS

D IALL 1-28

D IBIB ED ABS HITIND 29-40

D IALL 41-63

FILE 'HOME' ENTERED AT 13:10:34 ON 03 FEB 2006

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